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ATTENTION: APPLICATION BRANCH

Sir:

Transmitted herewith for filing is the patent application of

Inventor(s): Jean-Baptiste Dumas Milne Edwards, Aymeric Duclert, and Lydie Bougueleret

For: EXTENDED cDNAs

Enclosed are:

- (X) Sequence Listing in 392 pages.
- (X) Sequence Submission in 1 page.
- (X) Sequence Listing in computer readable form.
- (X) Fifteen (15) sheet(s) of drawing.
- (X) Return prepaid postcard.

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CLAIMS AS FILED

FOR	NUMBER FILED	NUMBER EXTRA	RATE	FEE
Basic Fee			\$760	\$760
Total Claims	20 - 20 =	0 ×	\$18	\$0
Independent Claims	17 - 3 =	14 ×	\$78	\$1,092
If application contains any multiple dependent claims(s), then add			\$260	\$0
TOTAL FILING FEE		\$1,852		

- (X) A check in the amount of \$1,852 to cover the filing fee is enclosed.
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*Daniel Hart*

Daniel Hart

Registration No. 40,637

Attorney of Record

**EXTENDED cDNAs**Related Applications

5 The present application claims priority from United States Provisional Patent Application Serial No. 60/069,957 filed December 17, 1997, United States Provisional Patent Application Serial No. 60/074,121 filed February 9, 1998, United States Provisional Patent Application Serial No. 60/081,563, filed April 13, 1998, United States Provisional Patent Application Serial No. 60/096,116, filed August 10, 1998, and United States Provisional Patent Application Serial No. 60/099,273 filed September 4, 1998 the  
10 disclosures of which are incorporated herein by reference in their entirety.

Table I lists the SEQ ID Nos. of the extended cDNAs in the present application, the SEQ ID Nos. of the extended cDNAs in the provisional applications, and the identities of the provisional applications in which the extended cDNAs were disclosed.

Background of the Invention

15 The estimated 50,000-100,000 genes scattered along the human chromosomes offer tremendous promise for the understanding, diagnosis, and treatment of human diseases. In addition, probes capable of specifically hybridizing to loci distributed throughout the human genome find applications in the construction of high resolution chromosome maps and in the identification of individuals.

20 In the past, the characterization of even a single human gene was a painstaking process, requiring years of effort. Recent developments in the areas of cloning vectors, DNA sequencing, and computer technology have merged to greatly accelerate the rate at which human genes can be isolated, sequenced, mapped, and characterized. Cloning vectors such as yeast artificial chromosomes (YACs) and bacterial artificial chromosomes  
25 (BACs) are able to accept DNA inserts ranging from 300 to 1000 kilobases (kb) or 100-400 kb in length respectively, thereby facilitating the manipulation and ordering of DNA sequences distributed over great distances on the human chromosomes. Automated DNA sequencing machines permit the rapid sequencing of human genes. Bioinformatics

software enables the comparison of nucleic acid and protein sequences, thereby assisting in the characterization of human gene products.

Currently, two different approaches are being pursued for identifying and characterizing the genes distributed along the human genome. In one approach, large fragments of genomic DNA are isolated, cloned, and sequenced. Potential open reading frames in these genomic sequences are identified using bio-informatics software. However, this approach entails sequencing large stretches of human DNA which do not encode proteins in order to find the protein encoding sequences scattered throughout the genome. In addition to requiring extensive sequencing, the bio-informatics software may mischaracterize the genomic sequences obtained. Thus, the software may produce false positives in which non-coding DNA is mischaracterized as coding DNA or false negatives in which coding DNA is mislabeled as non-coding DNA.

An alternative approach takes a more direct route to identifying and characterizing human genes. In this approach, complementary DNAs (cDNAs) are synthesized from isolated messenger RNAs (mRNAs) which encode human proteins. Using this approach, sequencing is only performed on DNA which is derived from protein coding portions of the genome. Often, only short stretches of the cDNAs are sequenced to obtain sequences called expressed sequence tags (ESTs). The ESTs may then be used to isolate or purify extended cDNAs which include sequences adjacent to the EST sequences. The extended cDNAs may contain all of the sequence of the EST which was used to obtain them or only a portion of the sequence of the EST which was used to obtain them. In addition, the extended cDNAs may contain the full coding sequence of the gene from which the EST was derived or, alternatively, the extended cDNAs may include portions of the coding sequence of the gene from which the EST was derived. It will be appreciated that there may be several extended cDNAs which include the EST sequence as a result of alternate splicing or the activity of alternative promoters.

In the past, the short EST sequences which could be used to isolate or purify extended cDNAs were often obtained from oligo-dT primed cDNA libraries. Accordingly, they mainly corresponded to the 3' untranslated region of the mRNA. In

part, the prevalence of EST sequences derived from the 3' end of the mRNA is a result of the fact that typical techniques for obtaining cDNAs, are not well suited for isolating cDNA sequences derived from the 5' ends of mRNAs. (Adams et al., *Nature* 377:174, 1996, Hillier et al., *Genome Res.* 6:807-828, 1996).

5           In addition, in those reported instances where longer cDNA sequences have been obtained, the reported sequences typically correspond to coding sequences and do not include the full 5' untranslated region of the mRNA from which the cDNA is derived. Such incomplete sequences may not include the first exon of the mRNA, particularly in situations where the first exon is short. Furthermore, they may not include some exons,  
10           often short ones, which are located upstream of splicing sites. Thus, there is a need to obtain sequences derived from the 5' ends of mRNAs which can be used to obtain extended cDNAs which may include the 5' sequences contained in the 5' ESTs.

          While many sequences derived from human chromosomes have practical applications, approaches based on the identification and characterization of those  
15           chromosomal sequences which encode a protein product are particularly relevant to diagnostic and therapeutic uses. Of the 50,000-100,000 protein coding genes, those genes encoding proteins which are secreted from the cell in which they are synthesized, as well as the secreted proteins themselves, are particularly valuable as potential therapeutic agents. Such proteins are often involved in cell to cell communication and may be  
20           responsible for producing a clinically relevant response in their target cells.

          In fact, several secretory proteins, including tissue plasminogen activator, G-CSF, GM-CSF, erythropoietin, human growth hormone, insulin, interferon- $\alpha$ , interferon- $\beta$ , interferon- $\gamma$ , and interleukin-2, are currently in clinical use. These proteins are used to treat a wide range of conditions, including acute myocardial infarction, acute ischemic  
25           stroke, anemia, diabetes, growth hormone deficiency, hepatitis, kidney carcinoma, chemotherapy induced neutropenia and multiple sclerosis. For these reasons, extended cDNAs encoding secreted proteins or portions thereof represent a particularly valuable source of therapeutic agents. Thus, there is a need for the identification and characterization of secreted proteins and the nucleic acids encoding them.



In addition to being therapeutically useful themselves, secretory proteins include short peptides, called signal peptides, at their amino termini which direct their secretion. These signal peptides are encoded by the signal sequences located at the 5' ends of the coding sequences of genes encoding secreted proteins. Because these signal peptides will direct the extracellular secretion of any protein to which they are operably linked, the signal sequences may be exploited to direct the efficient secretion of any protein by operably linking the signal sequences to a gene encoding the protein for which secretion is desired. This may prove beneficial in gene therapy strategies in which it is desired to deliver a particular gene product to cells other than the cell in which it is produced. Signal sequences encoding signal peptides also find application in simplifying protein purification techniques. In such applications, the extracellular secretion of the desired protein greatly facilitates purification by reducing the number of undesired proteins from which the desired protein must be selected. Thus, there exists a need to identify and characterize the 5' portions of the genes for secretory proteins which encode signal peptides.

Public information on the number of human genes for which the promoters and upstream regulatory regions have been identified and characterized is quite limited. In part, this may be due to the difficulty of isolating such regulatory sequences. Upstream regulatory sequences such as transcription factor binding sites are typically too short to be utilized as probes for isolating promoters from human genomic libraries. Recently, some approaches have been developed to isolate human promoters. One of them consists of making a CpG island library (Cross, S.H. et al., Purification of CpG Islands using a Methylated DNA Binding Column, *Nature Genetics* 6: 236-244 (1994)). The second consists of isolating human genomic DNA sequences containing SpeI binding sites by the use of SpeI binding protein. (Mortlock et al., *Genome Res.* 6:327-335, 1996). Both of these approaches have their limits due to a lack of specificity or of comprehensiveness.

5' ESTs and extended cDNAs obtainable therefrom may be used to efficiently identify and isolate upstream regulatory regions which control the location, developmental stage, rate, and quantity of protein synthesis, as well as the stability of the mRNA. (Theil

*et al.*, *BioFactors* 4:87-93, (1993). Once identified and characterized, these regulatory regions may be utilized in gene therapy or protein purification schemes to obtain the desired amount and locations of protein synthesis or to inhibit, reduce, or prevent the synthesis of undesirable gene products.

5 In addition, ESTs containing the 5' ends of secretory protein genes or extended cDNAs which include sequences adjacent to the sequences of the ESTs may include sequences useful as probes for chromosome mapping and the identification of individuals. Thus, there is a need to identify and characterize the sequences upstream of the 5' coding sequences of genes encoding secretory proteins.

#### 10 Summary of the Invention

The present invention relates to purified, isolated, or recombinant extended cDNAs which encode secreted proteins or fragments thereof. Preferably, the purified, isolated or recombinant cDNAs contain the entire open reading frame of their corresponding mRNAs, including a start codon and a stop codon. For example, the  
15 extended cDNAs may include nucleic acids encoding the signal peptide as well as the mature protein. Alternatively, the extended cDNAs may contain a fragment of the open reading frame. In some embodiments, the fragment may encode only the sequence of the mature protein. Alternatively, the fragment may encode only a portion of the mature protein. A further aspect of the present invention is a nucleic acid which encodes the  
20 signal peptide of a secreted protein.

The present extended cDNAs were obtained using ESTs which include sequences derived from the authentic 5' ends of their corresponding mRNAs. As used herein the terms "EST" or "5' EST" refer to the short cDNAs which were used to obtain the extended cDNAs of the present invention. As used herein, the term "extended cDNA" refers to the  
25 cDNAs which include sequences adjacent to the 5' EST used to obtain them. The extended cDNAs may contain all or a portion of the sequence of the EST which was used to obtain them. The term "corresponding mRNA" refers to the mRNA which was the template for the cDNA synthesis which produced the 5' EST. As used herein, the term "purified" does not require absolute purity; rather, it is intended as a relative definition.

Individual extended cDNA clones isolated from a cDNA library have been conventionally purified to electrophoretic homogeneity. The sequences obtained from these clones could not be obtained directly either from the library or from total human DNA. The extended cDNA clones are not naturally occurring as such, but rather are obtained via manipulation of a partially purified naturally occurring substance (messenger RNA). The conversion of mRNA into a cDNA library involves the creation of a synthetic substance (cDNA) and pure individual cDNA clones can be isolated from the synthetic library by clonal selection. Thus, creating a cDNA library from messenger RNA and subsequently isolating individual clones from that library results in an approximately  $10^4$ - $10^6$  fold purification of the native message. Purification of starting material or natural material to at least one order of magnitude, preferably two or three orders, and more preferably four or five orders of magnitude is expressly contemplated.

As used herein, the term "isolated" requires that the material be removed from its original environment (e.g., the natural environment if it is naturally occurring). For example, a naturally-occurring polynucleotide present in a living animal is not isolated, but the same polynucleotide, separated from some or all of the coexisting materials in the natural system, is isolated.

As used herein, the term "recombinant" means that the extended cDNA is adjacent to "backbone" nucleic acid to which it is not adjacent in its natural environment. Additionally, to be "enriched" the extended cDNAs will represent 5% or more of the number of nucleic acid inserts in a population of nucleic acid backbone molecules. Backbone molecules according to the present invention include nucleic acids such as expression vectors, self-replicating nucleic acids, viruses, integrating nucleic acids, and other vectors or nucleic acids used to maintain or manipulate a nucleic acid insert of interest. Preferably, the enriched extended cDNAs represent 15% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. More preferably, the enriched extended cDNAs represent 50% or more of the number of nucleic acid inserts in the population of recombinant backbone molecules. In a highly preferred embodiment, the enriched extended cDNAs represent 90% or more of the number of

nucleic acid inserts in the population of recombinant backbone molecules. "Stringent", "moderate," and "low" hybridization conditions are as defined in Example 29.

Unless otherwise indicated, a "complementary" sequence is fully complementary. Thus, extended cDNAs encoding secreted polypeptides or fragments thereof which are present in cDNA libraries in which one or more extended cDNAs encoding secreted polypeptides or fragments thereof make up 5% or more of the number of nucleic acid inserts in the backbone molecules are "enriched recombinant extended cDNAs" as defined herein. Likewise, extended cDNAs encoding secreted polypeptides or fragments thereof which are in a population of plasmids in which one or more extended cDNAs of the present invention have been inserted such that they represent 5% or more of the number of inserts in the plasmid backbone are "enriched recombinant extended cDNAs" as defined herein. However, extended cDNAs encoding secreted polypeptides or fragments thereof which are in cDNA libraries in which the extended cDNAs encoding secreted polypeptides or fragments thereof constitute less than 5% of the number of nucleic acid inserts in the population of backbone molecules, such as libraries in which backbone molecules having a cDNA insert encoding a secreted polypeptide are extremely rare, are not "enriched recombinant extended cDNAs."

In particular, the present invention relates to extended cDNAs which were derived from genes encoding secreted proteins. As used herein, a "secreted" protein is one which, when expressed in a suitable host cell, is transported across or through a membrane, including transport as a result of signal peptides in its amino acid sequence. "Secreted" proteins include without limitation proteins secreted wholly (e.g. soluble proteins), or partially (e.g. receptors) from the cell in which they are expressed. "Secreted" proteins also include without limitation proteins which are transported across the membrane of the endoplasmic reticulum.

Extended cDNAs encoding secreted proteins may include nucleic acid sequences, called signal sequences, which encode signal peptides which direct the extracellular secretion of the proteins encoded by the extended cDNAs. Generally, the signal peptides are located at the amino termini of secreted proteins.

Secreted proteins are translated by ribosomes associated with the "rough" endoplasmic reticulum. Generally, secreted proteins are co-translationally transferred to the membrane of the endoplasmic reticulum. Association of the ribosome with the endoplasmic reticulum during translation of secreted proteins is mediated by the signal peptide. The signal peptide is typically cleaved following its co-translational entry into the endoplasmic reticulum. After delivery to the endoplasmic reticulum, secreted proteins may proceed through the Golgi apparatus. In the Golgi apparatus, the proteins may undergo post-translational modification before entering secretory vesicles which transport them across the cell membrane.

The extended cDNAs of the present invention have several important applications. For example, they may be used to express the entire secreted protein which they encode. Alternatively, they may be used to express portions of the secreted protein. The portions may comprise the signal peptides encoded by the extended cDNAs or the mature proteins encoded by the extended cDNAs (i.e. the proteins generated when the signal peptide is cleaved off). The portions may also comprise polypeptides having at least 10 consecutive amino acids encoded by the extended cDNAs. Alternatively, the portions may comprise at least 15 consecutive amino acids encoded by the extended cDNAs. In some embodiments, the portions may comprise at least 25 consecutive amino acids encoded by the extended cDNAs. In other embodiments, the portions may comprise at least 40 amino acids encoded by the extended cDNAs.

Antibodies which specifically recognize the entire secreted proteins encoded by the extended cDNAs or fragments thereof having at least 10 consecutive amino acids, at least 15 consecutive amino acids, at least 25 consecutive amino acids, or at least 40 consecutive amino acids may also be obtained as described below. Antibodies which specifically recognize the mature protein generated when the signal peptide is cleaved may also be obtained as described below. Similarly, antibodies which specifically recognize the signal peptides encoded by the extended cDNAs may also be obtained.

In some embodiments, the extended cDNAs include the signal sequence. In other embodiments, the extended cDNAs may include the full coding sequence for the mature

protein (i.e. the protein generated when the signal polypeptide is cleaved off). In addition, the extended cDNAs may include regulatory regions upstream of the translation start site or downstream of the stop codon which control the amount, location, or developmental stage of gene expression. As discussed above, secreted proteins are therapeutically important. Thus, the proteins expressed from the cDNAs may be useful in treating or controlling a variety of human conditions. The extended cDNAs may also be used to obtain the corresponding genomic DNA. The term "corresponding genomic DNA" refers to the genomic DNA which encodes mRNA which includes the sequence of one of the strands of the extended cDNA in which thymidine residues in the sequence of the extended cDNA are replaced by uracil residues in the mRNA.

The extended cDNAs or genomic DNAs obtained therefrom may be used in forensic procedures to identify individuals or in diagnostic procedures to identify individuals having genetic diseases resulting from abnormal expression of the genes corresponding to the extended cDNAs. In addition, the present invention is useful for constructing a high resolution map of the human chromosomes.

The present invention also relates to secretion vectors capable of directing the secretion of a protein of interest. Such vectors may be used in gene therapy strategies in which it is desired to produce a gene product in one cell which is to be delivered to another location in the body. Secretion vectors may also facilitate the purification of desired proteins.

The present invention also relates to expression vectors capable of directing the expression of an inserted gene in a desired spatial or temporal manner or at a desired level. Such vectors may include sequences upstream of the extended cDNAs such as promoters or upstream regulatory sequences.

In addition, the present invention may also be used for gene therapy to control or treat genetic diseases. Signal peptides may also be fused to heterologous proteins to direct their extracellular secretion.

One embodiment of the present invention is a purified or isolated nucleic acid comprising the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence

complementary thereto. In one aspect of this embodiment, the nucleic acid is recombinant.

Another embodiment of the present invention is a purified or isolated nucleic acid comprising at least 10 consecutive bases of the sequence of one of SEQ ID NOs: 40-140 and 242-377 or one of the sequences complementary thereto. In one aspect of this embodiment, the nucleic acid comprises at least 15, 25, 30, 40, 50, 75, or 100 consecutive bases of one of the sequences of SEQ ID NOs: 40-140 and 242-377 or one of the sequences complementary thereto. The nucleic acid may be a recombinant nucleic acid.

Another embodiment of the present invention is a purified or isolated nucleic acid of at least 15 bases capable of hybridizing under stringent conditions to the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary to one of the sequences of SEQ ID NOs: 40-140 and 242-377. In one aspect of this embodiment, the nucleic acid is recombinant.

Another embodiment of the present invention is a purified or isolated nucleic acid comprising the full coding sequences of one of SEQ ID NOs: 40-140 and 242-377, wherein the full coding sequence optionally comprises the sequence encoding signal peptide as well as the sequence encoding mature protein. In a preferred embodiment, the isolated or purified nucleic acid comprises the full coding sequence of one of SEQ ID Nos. 40, 42-44, 46, 48, 49, 51, 53, 60, 62-72, 76-78, 80-83, 85-88, 90, 93, 94, 97, 99-102, 104, 107-125, 127, 132, 135-138, 140 and 242-377 wherein the full coding sequence comprises the sequence encoding signal peptide and the sequence encoding mature protein. In one aspect of this embodiment, the nucleic acid is recombinant.

A further embodiment of the present invention is a purified or isolated nucleic acid comprising the nucleotides of one of SEQ ID NOs: 40-140 and 242-377 which encode a mature protein. In a preferred embodiment, the purified or isolated nucleic acid comprises the nucleotides of one of SEQ ID NOs: 40-44, 46, 48, 49, 51-53, 55, 56, 58-72, 75-78, 80-88, 90, 93, 94, 97, 99-125, 127, 132, 133, 135-138, 140, and 242-377 which encode a mature protein. In one aspect of this embodiment, the nucleic acid is recombinant.

Yet another embodiment of the present invention is a purified or isolated nucleic acid comprising the nucleotides of one of SEQ ID NOs: 40-140 and 242-377 which encode the signal peptide. In a preferred embodiment, the purified or isolated nucleic acid comprises the nucleotides of SEQ ID NOs: 40, 42-46, 48, 49, 51, 53, 57, 60, 62-73, 76-78, 80-83, 85-88, 90, 93-95, 97, 99-102, 104, 107-125, 127, 128, 130, 132, 134-140 and 242-377 which encode the signal peptide. In one aspect of this embodiment, the nucleic acid is recombinant.

Another embodiment of the present invention is a purified or isolated nucleic acid encoding a polypeptide having the sequence of one of the sequences of SEQ ID NOs: 141-241 and 378-513.

Another embodiment of the present invention is a purified or isolated nucleic acid encoding a polypeptide having the sequence of a mature protein included in one of the sequences of SEQ ID NOs: 141-241 and 378-513. In a preferred embodiment, the purified or isolated nucleic acid encodes a polypeptide having the sequence of a mature protein included in one of the sequences of SEQ ID NOs: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513.

Another embodiment of the present invention is a purified or isolated nucleic acid encoding a polypeptide having the sequence of a signal peptide included in one of the sequences of SEQ ID NOs: 141-241 and 378-513. In a preferred embodiment, the purified or isolated nucleic acid encodes a polypeptide having the sequence of a signal peptide included in one of the sequences of SEQ ID NOs: 141, 143-147, 149, 150, 152, 154, 158, 161, 163-174, 177-179, 181-184, 186-189, 191, 194-196, 198, 200-203, 205, 208-226, 228, 229, 231, 233, 235-241, and 378-513.

Yet another embodiment of the present invention is a purified or isolated protein comprising the sequence of one of SEQ ID NOs: 141-241 and 378-513.

Another embodiment of the present invention is a purified or isolated polypeptide comprising at least 10 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In one aspect of this embodiment, the purified or isolated



polypeptide comprises at least 15, 20, 25, 35, 50, 75, 100, 150 or 200 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In still another aspect, the purified or isolated polypeptide comprises at least 25 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513.

5 Another embodiment of the present invention is an isolated or purified polypeptide comprising a signal peptide of one of the polypeptides of SEQ ID NOs: 141-241 and 378-513. In a preferred embodiment, the isolated or purified polypeptide comprises a signal peptide of one of the polypeptides of SEQ ID NOs: 141, 143-147, 149, 150, 152, 154, 158, 161, 163-174, 177-179, 181-184, 186-189, 191, 194-196, 198, 200-203, 205, 208-10 226, 228, 229, 231, 233, 235-241, and 378-513.

Yet another embodiment of the present invention is an isolated or purified polypeptide comprising a mature protein of one of the polypeptides of SEQ ID NOs: 141-241 and 378-513. In a preferred embodiment, the isolated or purified polypeptide comprises a mature protein of one of the polypeptides of SEQ ID NOs: 141-145, 147, 15 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513.

A further embodiment of the present invention is a method of making a protein comprising one of the sequences of SEQ ID NO: 141-241 and 378-513, comprising the steps of obtaining a cDNA comprising one of the sequences of sequence of SEQ ID NO: 20 40-140 and 242-377, inserting the cDNA in an expression vector such that the cDNA is operably linked to a promoter, and introducing the expression vector into a host cell whereby the host cell produces the protein encoded by said cDNA. In one aspect of this embodiment, the method further comprises the step of isolating the protein.

Another embodiment of the present invention is a protein obtainable by the 25 method described in the preceding paragraph.

Another embodiment of the present invention is a method of making a protein comprising the amino acid sequence of the mature protein contained in one of the sequences of SEQ ID NO: 141-241 and 378-513, comprising the steps of obtaining a cDNA comprising one of the nucleotides sequence of sequence of SEQ ID NO: 40-140

and 242-377 which encode for the mature protein, inserting the cDNA in an expression vector such that the cDNA is operably linked to a promoter, and introducing the expression vector into a host cell whereby the host cell produces the mature protein encoded by the cDNA. In one aspect of this embodiment, the method further comprises the step of isolating the protein.

Another embodiment of the present invention is a mature protein obtainable by the method described in the preceding paragraph.

In a preferred embodiment, the above method comprises a method of making a protein comprising the amino acid sequence of the mature protein contained in one of the sequences of SEQ ID NO: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513, comprising the steps of obtaining a cDNA comprising one of the nucleotides sequence of sequence of SEQ ID NO: 40-44, 46, 48, 49, 51-53, 55, 56, 58-72, 75-78, 80-88, 90, 93, 94, 97, 99-125, 127, 132, 133, 135-138, 140, and 242-377 which encode for the mature protein, inserting the cDNA in an expression vector such that the cDNA is operably linked to a promoter, and introducing the expression vector into a host cell whereby the host cell produces the mature protein encoded by the cDNA. In one aspect of this embodiment, the method further comprises the step of isolating the protein.

Another embodiment of the present invention is a host cell containing the purified or isolated nucleic acids comprising the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary thereto described herein.

Another embodiment of the present invention is a host cell containing the purified or isolated nucleic acids comprising the full coding sequences of one of SEQ ID NOs: 40-140 and 242-377, wherein the full coding sequence comprises the sequence encoding signal peptide and the sequence encoding mature protein described herein.

Another embodiment of the present invention is a host cell containing the purified or isolated nucleic acids comprising the nucleotides of one of SEQ ID NOs: 40-140 and 242-377 which encode a mature protein which are described herein. Preferably, the host cell contains the purified or isolated nucleic acids comprising the nucleotides of one of

SEQ ID NOs: 40-44, 46, 48, 49, 51-53, 55, 56, 58-72, 75-78, 80-88, 90, 93, 94, 97, 99-125, 127, 132, 133, 135-138, 140, and 242-377 which encode a mature protein.

Another embodiment of the present invention is a host cell containing the purified or isolated nucleic acids comprising the nucleotides of one of SEQ ID NOs: 40-140 and 242-377 which encode the signal peptide which are described herein. Preferably, the host cell contains the purified or isolated nucleic acids comprising the nucleotides of one of SEQ ID Nos.: 40, 42-46, 48, 49, 51, 53, 57, 60, 62-73, 76-78, 80-83, 85-88, 90, 93-95, 97, 99-102, 104, 107-125, 127, 128, 130, 132, 134-140 and 242-377 which encode the signal peptide.

Another embodiment of the present invention is a purified or isolated antibody capable of specifically binding to a protein having the sequence of one of SEQ ID NOs: 141-241 and 378-513. In one aspect of this embodiment, the antibody is capable of binding to a polypeptide comprising at least 10 consecutive amino acids of the sequence of one of SEQ ID NOs: 141-241 and 378-513.

Another embodiment of the present invention is an array of cDNAs or fragments thereof of at least 15 nucleotides in length which includes at least one of the sequences of SEQ ID NOs: 40-140 and 242-377, or one of the sequences complementary to the sequences of SEQ ID NOs: 40-140 and 242-377, or a fragment thereof of at least 15 consecutive nucleotides. In one aspect of this embodiment, the array includes at least two of the sequences of SEQ ID NOs: 40-140 and 242-377, the sequences complementary to the sequences of SEQ ID NOs: 40-140 and 242-377, or fragments thereof of at least 15 consecutive nucleotides. In another aspect of this embodiment, the array includes at least five of the sequences of SEQ ID NOs: 40-140 and 242-377, the sequences complementary to the sequences of SEQ ID NOs: 40-140 and 242-377, or fragments thereof of at least 15 consecutive nucleotides.

A further embodiment of the invention encompasses purified polynucleotides comprising an insert from a clone deposited in a deposit having an accession number selected from the group consisting of the accession numbers listed in Table VI or a fragment thereof comprising a contiguous span of at least 8, 10, 12, 15, 20, 25, 40, 60,

100, or 200 nucleotides of said insert. An additional embodiment of the invention encompasses purified polypeptides which comprise, consist of, or consist essentially of an amino acid sequence encoded by the insert from a clone deposited in a deposit having an accession number selected from the group consisting of the accession numbers listed in  
5 Table VI, as well as polypeptides which comprise a fragment of said amino acid sequence consisting of a signal peptide, a mature protein, or a contiguous span of at least 5, 8, 10, 12, 15, 20, 25, 40, 60, 100, or 200 amino acids encoded by said insert.

An additional embodiment of the invention encompasses purified polypeptides which comprise a contiguous span of at least 5, 8, 10, 12, 15, 20, 25, 40, 60, 100, or 200  
10 amino acids of SEQ ID NOs: 158, 174, 175, 196, 226, 231, 232, wherein said contiguous span comprises at least one of the amino acid positions which was not shown to be identical to a public sequence in any of Figures 11 to 15. Also encompassed by the invention are purified polynucleotides encoding said polypeptides.

#### Brief Description of the Drawings

15 Figure 1 is a summary of a procedure for obtaining cDNAs which have been selected to include the 5' ends of the mRNAs from which they are derived.

Figure 2 is an analysis of the 43 amino terminal amino acids of all human SwissProt proteins to determine the frequency of false positives and false negatives using the techniques for signal peptide identification described herein.

20 Figure 3 shows the distribution of von Heijne scores for 5' ESTs in each of the categories described herein and the probability that these 5' ESTs encode a signal peptide.

Figure 4 shows the distribution of 5' ESTs in each category and the number of 5' ESTs in each category having a given minimum von Heijne's score.

25 Figure 5 shows the tissues from which the mRNAs corresponding to the 5' ESTs in each of the categories described herein were obtained.

Figure 6 illustrates a method for obtaining extended cDNAs.

Figure 7 is a map of pED6dpc2.

Figure 8 provides a schematic description of the promoters isolated and the way they are assembled with the corresponding 5' tags.

Figure 9 describes the transcription factor binding sites present in each of these promoters.

Figure 10 is an alignment of the protein of SEQ ID NO: 217 with the human protein TFAR19 that may play a role in apoptosis (Genbank accession number AF014955, SEQ ID NO: 516).

Figure 11 is an alignment of the proteins of SEQ ID NOs: 174, 175 and 232 with a human secreted protein (Genseq accession number W36955, SEQ ID NO: 517).

Figure 12 is an alignment of the protein of SEQ ID NO: 231 with the human E25 protein (Genbank accession number AF038953, SEQ ID NO: 515).

Figure 13 is an alignment of the protein of SEQ ID NO: 196 with the human seventransmembrane protein (Genbank accession number Y11395, SEQ ID NO: 518).

Figure 14 is an alignment of the protein of SEQ ID NOs: 158 with the murine subunit 7a of the COP9 complex (Genbank accession number AF071316, SEQ ID NO: 519).

Figure 15 is an alignment of the protein of SEQ ID NO: 226 with the bovine subunit B14.5B of the NADH-ubiquinone oxidoreductase complex (Arizmendi *et al*, *FEBS Lett.*, **313** : 80-84 (1992) and Swissprot accession -number Q02827, SEQ ID NO: 514).

### Detailed Description of the Preferred Embodiment

#### **I. Obtaining 5' ESTs**

The present extended cDNAs were obtained using 5' ESTs which were isolated as described below.

##### A. Chemical Methods for Obtaining mRNAs having Intact 5' Ends

In order to obtain the 5' ESTs used to obtain the extended cDNAs of the present invention, mRNAs having intact 5' ends must be obtained. Currently, there are two approaches for obtaining such mRNAs. One of these approaches is a chemical modification method involving derivatization of the 5' ends of the mRNAs and selection of the derivatized mRNAs. The 5' ends of eucaryotic mRNAs possess a structure referred to as a "cap" which comprises a guanosine methylated at the 7 position. The cap is joined

to the first transcribed base of the mRNA by a 5', 5'-triphosphate bond. In some instances, the 5' guanosine is methylated in both the 2 and 7 positions. Rarely, the 5' guanosine is trimethylated at the 2, 7 and 7 positions. In the chemical method for obtaining mRNAs having intact 5' ends, the 5' cap is specifically derivatized and coupled to a reactive group on an immobilizing substrate. This specific derivatization is based on the fact that only the ribose linked to the methylated guanosine at the 5' end of the mRNA and the ribose linked to the base at the 3' terminus of the mRNA, possess 2', 3'-cis diols. Optionally, where the 3' terminal ribose has a 2', 3'-cis diol, the 2', 3'-cis diol at the 3' end may be chemically modified, substituted, converted, or eliminated, leaving only the ribose linked to the methylated guanosine at the 5' end of the mRNA with a 2', 3'-cis diol. A variety of techniques are available for eliminating the 2', 3'-cis diol on the 3' terminal ribose. For example, controlled alkaline hydrolysis may be used to generate mRNA fragments in which the 3' terminal ribose is a 3'-phosphate, 2'-phosphate or (2', 3')-cyclophosphate. Thereafter, the fragment which includes the original 3' ribose may be eliminated from the mixture through chromatography on an oligo-dT column. Alternatively, a base which lacks the 2', 3'-cis diol may be added to the 3' end of the mRNA using an RNA ligase such as T4 RNA ligase. Example 1 below describes a method for ligation of pCp to the 3' end of messenger RNA.

### EXAMPLE 1

#### Ligation of the Nucleoside Diphosphate pCp to the 3' End of Messenger RNA

1 µg of RNA was incubated in a final reaction medium of 10 µl in the presence of 5 U of T<sub>4</sub> phage RNA ligase in the buffer provided by the manufacturer (Gibco - BRL), 40 U of the RNase inhibitor RNasin (Promega) and, 2 µl of <sup>32</sup>pCp (Amersham #PB 10208).

The incubation was performed at 37°C for 2 hours or overnight at 7-8°C.

Following modification or elimination of the 2', 3'-cis diol at the 3' ribose, the 2', 3'-cis diol present at the 5' end of the mRNA may be oxidized using reagents such as NaBH<sub>4</sub>, NaBH<sub>3</sub>CN, or sodium periodate, thereby converting the 2', 3'-cis diol to a dialdehyde. Example 2 describes the oxidation of the 2', 3'-cis diol at the 5' end of the mRNA with sodium periodate.

## EXAMPLE 2

### Oxidation of 2', 3'-cis diol at the 5' End of the mRNA

0.1 OD unit of either a capped oligoribonucleotide of 47 nucleotides (including the cap) or an uncapped oligoribonucleotide of 46 nucleotides were treated as follows. The oligoribonucleotides were produced by in vitro transcription using the transcription kit "AmpliScribe T7" (Epicentre Technologies). As indicated below, the DNA template for the RNA transcript contained a single cytosine. To synthesize the uncapped RNA, all four NTPs were included in the in vitro transcription reaction. To obtain the capped RNA, GTP was replaced by an analogue of the cap, m7G(5')ppp(5')G. This compound, recognized by polymerase, was incorporated into the 5' end of the nascent transcript during the step of initiation of transcription but was not capable of incorporation during the extension step. Consequently, the resulting RNA contained a cap at its 5' end. The sequences of the oligoribonucleotides produced by the in vitro transcription reaction were:

+Cap:

5'm7GpppGCAUCCUACUCCCAUCCAAUUCCACCCUAACUCCUCCCAUCUCCAC-3' (SEQ ID NO:1)

-Cap:

5'-pppGCAUCCUACUCCCAUCCAAUUCCACCCUAACUCCUCCCAUCUCCAC-3' (SEQ ID NO:2)

The oligoribonucleotides were dissolved in 9 µl of acetate buffer (0.1 M sodium acetate, pH 5.2) and 3 µl of freshly prepared 0.1 M sodium periodate solution. The mixture was incubated for 1 hour in the dark at 4°C or room temperature. Thereafter, the reaction was stopped by adding 4 µl of 10% ethylene glycol. The product was ethanol precipitated, resuspended in 10µl or more of water or appropriate buffer and dialyzed against water.

The resulting aldehyde groups may then be coupled to molecules having a reactive amine group, such as hydrazine, carbazide, thiocarbazide or semicarbazide groups, in order to facilitate enrichment of the 5' ends of the mRNAs. Molecules having reactive amine groups which are suitable for use in selecting mRNAs having intact 5' ends include

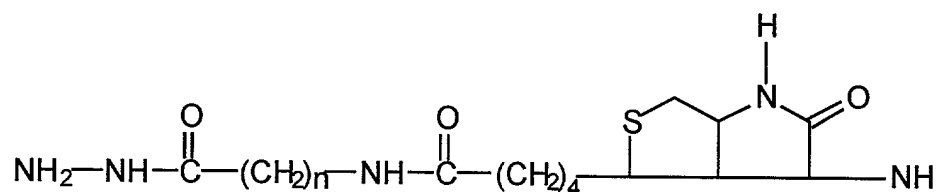
avidin, proteins, antibodies, vitamins, ligands capable of specifically binding to receptor molecules, or oligonucleotides. Example 3 below describes the coupling of the resulting dialdehyde to biotin.

### EXAMPLE 3

5

#### Coupling of the Dialdehyde with Biotin

The oxidation product obtained in Example 2 was dissolved in 50  $\mu$ l of sodium acetate at a pH of between 5 and 5.2 and 50  $\mu$ l of freshly prepared 0.02 M solution of biotin hydrazide in a methoxyethanol/watermixture (1:1) of formula:



10

In the compound used in these experiments,  $n=5$ . However, it will be appreciated that other commercially available hydrazides may also be used, such as molecules of the formula above in which  $n$  varies from 0 to 5.

The mixture was then incubated for 2 hours at 37°C. Following the incubation, the mixture was precipitated with ethanol and dialyzed against distilled water.

15

Example 4 demonstrates the specificity of the biotinylation reaction.

### EXAMPLE 4

#### Specificity of Biotinylation

The specificity of the biotinylation for capped mRNAs was evaluated by gel electrophoresis of the following samples:

20

Sample 1. The 46 nucleotide uncapped in vitro transcript prepared as in Example 2 and labeled with  $^{32}$ pCp as described in Example 1.



Sample 2. The 46 nucleotide uncapped in vitro transcript prepared as in Example 2, labeled with  $^{32}\text{pCp}$  as described in Example 1, treated with the oxidation reaction of Example 2, and subjected to the biotinylation conditions of Example 3.

5 Sample 3. The 47 nucleotide capped in vitro transcript prepared as in Example 2 and labeled with  $^{32}\text{pCp}$  as described in Example 1.

Sample 4. The 47 nucleotide capped in vitro transcript prepared as in Example 2, labeled with  $^{32}\text{pCp}$  as described in Example 1, treated with the oxidation reaction of Example 2, and subjected to the biotinylation conditions of Example 3.

10 Samples 1 and 2 had identical migration rates, demonstrating that the uncapped RNAs were not oxidized and biotinylated. Sample 3 migrated more slowly than Samples 1 and 2, while Sample 4 exhibited the slowest migration. The difference in migration of the RNAs in Samples 3 and 4 demonstrates that the capped RNAs were specifically biotinylated.

15 In some cases, mRNAs having intact 5' ends may be enriched by binding the molecule containing a reactive amine group to a suitable solid phase substrate such as the inside of the vessel containing the mRNAs, magnetic beads, chromatography matrices, or nylon or nitrocellulose membranes. For example, where the molecule having a reactive amine group is biotin, the solid phase substrate may be coupled to avidin or streptavidin. Alternatively, where the molecule having the reactive amine group is an antibody or  
20 receptor ligand, the solid phase substrate may be coupled to the cognate antigen or receptor. Finally, where the molecule having a reactive amine group comprises an oligonucleotide, the solid phase substrate may comprise a complementary oligonucleotide.

25 The mRNAs having intact 5' ends may be released from the solid phase following the enrichment procedure. For example, where the dialdehyde is coupled to biotin hydrazide and the solid phase comprises streptavidin, the mRNAs may be released from the solid phase by simply heating to 95 degrees Celsius in 2% SDS. In some methods, the molecule having a reactive amine group may also be cleaved from the mRNAs having intact 5' ends following enrichment. Example 5 describes the capture of biotinylated

mRNAs with streptavidin coated beads and the release of the biotinylated mRNAs from the beads following enrichment.

### EXAMPLE 5

#### Capture and Release of Biotinylated mRNAs Using Streptavidin Coated Beads

5       The streptavidin-coated magnetic beads were prepared according to the manufacturer's instructions (CPG Inc., USA). The biotinylated mRNAs were added to a hybridization buffer (1.5 M NaCl, pH 5 - 6). After incubating for 30 minutes, the unbound and nonbiotinylated material was removed. The beads were washed several times in water with 1% SDS. The beads obtained were incubated for 15 minutes at 95°C in water  
10       containing 2% SDS.

Example 6 demonstrates the efficiency with which biotinylated mRNAs were recovered from the streptavidin coated beads.

### EXAMPLE 6

#### Efficiency of Recovery of Biotinylated mRNAs

15       The efficiency of the recovery procedure was evaluated as follows. RNAs were labeled with <sup>32</sup>pCp, oxidized, biotinylated and bound to streptavidin coated beads as described above. Subsequently, the bound RNAs were incubated for 5, 15 or 30 minutes at 95°C in the presence of 2% SDS.

20       The products of the reaction were analyzed by electrophoresis on 12% polyacrylamide gels under denaturing conditions (7 M urea). The gels were subjected to autoradiography. During this manipulation, the hydrazone bonds were not reduced.

Increasing amounts of nucleic acids were recovered as incubation times in 2% SDS increased, demonstrating that biotinylated mRNAs were efficiently recovered.

25       In an alternative method for obtaining mRNAs having intact 5' ends, an oligonucleotide which has been derivatized to contain a reactive amine group is specifically coupled to mRNAs having an intact cap. Preferably, the 3' end of the mRNA is blocked prior to the step in which the aldehyde groups are joined to the derivatized oligonucleotide, as described above, so as to prevent the derivatized oligonucleotide from being joined to the 3' end of the mRNA. For example, pCp may be attached to the 3' end

of the mRNA using T4 RNA ligase. However, as discussed above, blocking the 3' end of the mRNA is an optional step. Derivatized oligonucleotides may be prepared as described below in Example 7.

### EXAMPLE 7

5

#### Derivatization of the Oligonucleotide

An oligonucleotide phosphorylated at its 3' end was converted to a 3' hydrazide in 3' by treatment with an aqueous solution of hydrazine or of dihydrazide of the formula  $H_2N(R1)NH_2$  at about 1 to 3 M, and at pH 4.5, in the presence of a carbodiimide type agent soluble in water such as 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide at a final concentration of 0.3 M at a temperature of 8°C overnight.

10

The derivatized oligonucleotide was then separated from the other agents and products using a standard technique for isolating oligonucleotides.

15

As discussed above, the mRNAs to be enriched may be treated to eliminate the 3' OH groups which may be present thereon. This may be accomplished by enzymatic ligation of sequences lacking a 3' OH, such as pCp, as described above in Example 1. Alternatively, the 3' OH groups may be eliminated by alkaline hydrolysis as described in Example 8 below.

### EXAMPLE 8

#### Alkaline Hydrolysis of mRNA

20

The mRNAs may be treated with alkaline hydrolysis as follows. In a total volume of 100µl of 0.1N sodium hydroxide, 1.5µg mRNA is incubated for 40 to 60 minutes at 4°C. The solution is neutralized with acetic acid and precipitated with ethanol.

Following the optional elimination of the 3' OH groups, the diol groups at the 5' ends of the mRNAs are oxidized as described below in Example 9.

25

### EXAMPLE 9

#### Oxidation of Diols

Up to 1 OD unit of RNA was dissolved in 9 µl of buffer (0.1 M sodium acetate, pH 6-7 or water) and 3 µl of freshly prepared 0.1 M sodium periodate solution. The reaction was incubated for 1 h in the dark at 4°C or room temperature. Following the

incubation, the reaction was stopped by adding 4  $\mu$ l of 10% ethylene glycol. Thereafter the mixture was incubated at room temperature for 15 minutes. After ethanol precipitation, the product was resuspended in 10 $\mu$ l or more of water or appropriate buffer and dialyzed against water.

- 5        Following oxidation of the diol groups at the 5' ends of the mRNAs, the derivatized oligonucleotide was joined to the resulting aldehydes as described in Example 10.

### EXAMPLE 10

#### Reaction of Aldehydes with Derivatized Oligonucleotides

- 10        The oxidized mRNA was dissolved in an acidic medium such as 50  $\mu$ l of sodium acetate pH 4-6. 50  $\mu$ l of a solution of the derivatized oligonucleotide was added such that an mRNA:derivatized oligonucleotide ratio of 1:20 was obtained and mixture was reduced with a borohydride. The mixture was allowed to incubate for 2 h at 37°C or overnight (14 h) at 10°C. The mixture was ethanol precipitated, resuspended in 10 $\mu$ l or more of
- 15        water or appropriate buffer and dialyzed against distilled water. If desired, the resulting product may be analyzed using acrylamide gel electrophoresis, HPLC analysis, or other conventional techniques.

Following the attachment of the derivatized oligonucleotide to the mRNAs, a reverse transcription reaction may be performed as described in Example 11 below.

- 20        **EXAMPLE 11**

#### Reverse Transcription of mRNAs

- 25        An oligodeoxyribonucleotide was derivatized as follows. 3 OD units of an oligodeoxyribonucleotide of sequence ATCAAGAATTCGCACGAGACCATTA (SEQ ID NO:3) having 5'-OH and 3'-P ends were dissolved in 70  $\mu$ l of a 1.5 M hydroxybenzotriazole solution, pH 5.3, prepared in dimethylformamide/water (75:25) containing 2  $\mu$ g of 1-ethyl-3-(3-dimethylaminopropyl)carbodiimide. The mixture was incubated for 2 h 30 min at 22°C. The mixture was then precipitated twice in LiClO<sub>4</sub>/acetone. The pellet was resuspended in 200  $\mu$ l of 0.25 M hydrazine and incubated

at 8°C from 3 to 14 h. Following the hydrazine reaction, the mixture was precipitated twice in LiClO<sub>4</sub>/acetone.

5 The messenger RNAs to be reverse transcribed were extracted from blocks of placenta having sides of 2 cm which had been stored at -80°C. The mRNA was extracted using conventional acidic phenol techniques. Oligo-dT chromatography was used to purify the mRNAs. The integrity of the mRNAs was checked by Northern-blotting.

10 The diol groups on 7 µg of the placental mRNAs were oxidized as described above in Example 9. The derivatized oligonucleotide was joined to the mRNAs as described in Example 10 above except that the precipitation step was replaced by an exclusion chromatography step to remove derivatized oligodeoxyribonucleotides which were not joined to mRNAs. Exclusion chromatography was performed as follows:

15 10 ml of AcA34 (BioSeptra#230151) gel were equilibrated in 50 ml of a solution of 10 mM Tris pH 8.0, 300 mM NaCl, 1 mM EDTA, and 0.05% SDS. The mixture was allowed to sediment. The supernatant was eliminated and the gel was resuspended in 50 ml of buffer. This procedure was repeated 2 or 3 times.

A glass bead (diameter 3 mm) was introduced into a 2 ml disposable pipette (length 25 cm). The pipette was filled with the gel suspension until the height of the gel stabilized at 1 cm from the top of the pipette. The column was then equilibrated with 20 ml of equilibration buffer (10 mM Tris HCl pH 7.4, 20 mM NaCl).

20 10 µl of the mRNA which had been reacted with the derivatized oligonucleotide were mixed in 39 µl of 10 mM urea and 2 µl of blue-glycerol buffer, which had been prepared by dissolving 5 mg of bromophenol blue in 60% glycerol (v/v), and passing the mixture through a filter with a filter of diameter 0.45 µm.

25 The column was loaded. As soon as the sample had penetrated, equilibration buffer was added. 100 µl fractions were collected. Derivatized oligonucleotide which had not been attached to mRNA appeared in fraction 16 and later fractions. Fractions 3 to 15 were combined and precipitated with ethanol.

The mRNAs which had been reacted with the derivatized oligonucleotide were spotted on a nylon membrane and hybridized to a radioactive probe using conventional

techniques. The radioactive probe used in these hybridizations was an oligodeoxyribonucleotide of sequence TAATGGTCTCGTGCGAATTCTTGAT (SEQ ID NO:4) which was anticomplementary to the derivatized oligonucleotide and was labeled at its 5' end with  $^{32}\text{P}$ . 1/10th of the mRNAs which had been reacted with the derivatized oligonucleotide was spotted in two spots on the membrane and the membrane was visualized by autoradiography after hybridization of the probe. A signal was observed, indicating that the derivatized oligonucleotide had been joined to the mRNA.

The remaining 9/10 of the mRNAs which had been reacted with the derivatized oligonucleotide was reverse transcribed as follows. A reverse transcription reaction was carried out with reverse transcriptase following the manufacturer's instructions. To prime the reaction, 50 pmol of nonamers with random sequence were used.

A portion of the resulting cDNA was spotted on a positively charged nylon membrane using conventional methods. The cDNAs were spotted on the membrane after the cDNA:RNA heteroduplexes had been subjected to an alkaline hydrolysis in order to eliminate the RNAs. An oligonucleotide having a sequence identical to that of the derivatized oligonucleotide was labeled at its 5' end with  $^{32}\text{P}$  and hybridized to the cDNA blots using conventional techniques. Single-stranded cDNAs resulting from the reverse transcription reaction were spotted on the membrane. As controls, the blot contained 1 pmol, 100 fmol, 50 fmol, 10 fmol and 1 fmol respectively of a control oligodeoxyribonucleotide of sequence identical to that of the derivatized oligonucleotide. The signal observed in the spots containing the cDNA indicated that approximately 15 fmol of the derivatized oligonucleotide had been reverse transcribed.

These results demonstrate that the reverse transcription can be performed through the cap and, in particular, that reverse transcriptase crosses the 5'-P-P-P-5' bond of the cap of eukaryotic messenger RNAs.

The single stranded cDNAs obtained after the above first strand synthesis were used as template for PCR reactions. Two types of reactions were carried out. First, specific amplification of the mRNAs for the alpha globin, dehydrogenase, pp15 and

elongation factor E4 were carried out using the following pairs of oligodeoxyribonucleotideprimers.

alpha-globin

GLO-S: CCG ACA AGA CCA ACG TCA AGG CCG C (SEQ ID NO:5)

5 GLO-As: TCA CCA GCA GGC AGT GGC TTA GGA G 3' (SEQ ID NO:6)

dehydrogenase

3 DH-S: AGT GAT TCC TGC TAC TTT GGA TGG C (SEQ ID NO:7)

3 DH-As: GCT TGG TCT TGT TCT GGA GTT TAG A (SEQ ID NO:8)

pp15

10 PP15-S: TCC AGA ATG GGA GAC AAG CCA ATT T (SEQ ID NO:9)

PP15-As: AGG GAG GAG GAA ACA GCG TGA GTC C (SEQ ID NO:10)

Elongation factor E4

EFA1-S: ATG GGA AAG GAA AAG ACT CAT ATC A (SEQ ID NO:11)

EF1A-As: AGC AGC AAC AAT CAG GAC AGC ACA G (SEQ ID NO:12)

15 Non specific amplifications were also carried out with the antisense (\_As) oligodeoxyribonucleotides of the pairs described above and a primer chosen from the sequence of the derivatized oligodeoxyribonucleotide (ATCAAGAATTCGCACGAGACCATTA)(SEQ ID NO:13).

20 A 1.5% agarose gel containing the following samples corresponding to the PCR products of reverse transcription was stained with ethidium bromide. (1/20th of the products of reverse transcription were used for each PCR reaction).

Sample 1: The products of a PCR reaction using the globin primers of SEQ ID NOs 5 and 6 in the presence of cDNA.

25 Sample 2: The products of a PCR reaction using the globin primers of SEQ ID NOs 5 and 6 in the absence of added cDNA.

Sample 3: The products of a PCR reaction using the dehydrogenase primers of SEQ ID NOs 7 and 8 in the presence of cDNA.

Sample 4: The products of a PCR reaction using the dehydrogenase primers of SEQ ID NOs 7 and 8 in the absence of added cDNA.

Sample 5: The products of a PCR reaction using the pp15 primers of SEQ ID NOs 9 and 10 in the presence of cDNA.

Sample 6: The products of a PCR reaction using the pp15 primers of SEQ ID NOs 9 and 10 in the absence of added cDNA.

5        Sample 7: The products of a PCR reaction using the EIE4 primers of SEQ ID NOs 11 and 12 in the presence of added cDNA.

Sample 8: The products of a PCR reaction using the EIE4 primers of SEQ ID NOs 11 and 12 in the absence of added cDNA.

10        In Samples 1, 3, 5 and 7, a band of the size expected for the PCR product was observed, indicating the presence of the corresponding sequence in the cDNA population.

15        PCR reactions were also carried out with the antisense oligonucleotides of the globin and dehydrogenase primers (SEQ ID NOs 6 and 8) and an oligonucleotide whose sequence corresponds to that of the derivatized oligonucleotide. The presence of PCR products of the expected size in the samples corresponding to samples 1 and 3 above indicated that the derivatized oligonucleotide had been incorporated.

The above examples summarize the chemical procedure for enriching mRNAs for those having intact 5' ends. Further detail regarding the chemical approaches for obtaining mRNAs having intact 5' ends are disclosed in International Application No. WO96/34981, published November 7, 1996, which is incorporated herein by reference.

20        Strategies based on the above chemical modifications to the 5' cap structure may be utilized to generate cDNAs which have been selected to include the 5' ends of the mRNAs from which they are derived. In one version of such procedures, the 5' ends of the mRNAs are modified as described above. Thereafter, a reverse transcription reaction is conducted to extend a primer complementary to the mRNA to the 5' end of the mRNA.

25        Single stranded RNAs are eliminated to obtain a population of cDNA/mRNA heteroduplexes in which the mRNA includes an intact 5' end. The resulting heteroduplexes may be captured on a solid phase coated with a molecule capable of interacting with the molecule used to derivatize the 5' end of the mRNA. Thereafter, the strands of the heteroduplexes are separated to recover single stranded first cDNA strands



which include the 5' end of the mRNA. Second strand cDNA synthesis may then proceed using conventional techniques. For example, the procedures disclosed in WO 96/34981 or in Carninci, P. et al. High-Efficiency Full-Length cDNA Cloning by Biotinylated CAP Trapper. **Genomics** 37:327-336 (1996), the disclosures of which are incorporated herein  
5 by reference, may be employed to select cDNAs which include the sequence derived from the 5' end of the coding sequence of the mRNA.

Following ligation of the oligonucleotide tag to the 5' cap of the mRNA, a reverse transcription reaction is conducted to extend a primer complementary to the mRNA to the 5' end of the mRNA. Following elimination of the RNA component of the resulting  
10 heteroduplex using standard techniques, second strand cDNA synthesis is conducted with a primer complementary to the oligonucleotide tag.

Figure 1 summarizes the above procedures for obtaining cDNAs which have been selected to include the 5' ends of the mRNAs from which they are derived.

#### B. Enzymatic Methods for Obtaining mRNAs having Intact 5' Ends

15 Other techniques for selecting cDNAs extending to the 5' end of the mRNA from which they are derived are fully enzymatic. Some versions of these techniques are disclosed in Dumas Milne Edwards J.B. (Doctoral Thesis of Paris VI University, Le clonage des ADNc complets: difficultes et perspectives nouvelles. Apports pour l'etude de la regulation de l'expression de la tryptophane hydroxylase de rat, 20 Dec. 1993), EP0  
20 625572 and Kato et al. Construction of a Human Full-Length cDNA Bank. *Gene* 150:243-250 (1994), the disclosures of which are incorporated herein by reference.

Briefly, in such approaches, isolated mRNA is treated with alkaline phosphatase to remove the phosphate groups present on the 5' ends of uncapped incomplete mRNAs. Following this procedure, the cap present on full length mRNAs is enzymatically removed  
25 with a decapping enzyme such as T4 polynucleotide kinase or tobacco acid pyrophosphatase. An oligonucleotide, which may be either a DNA oligonucleotide or a DNA-RNA hybrid oligonucleotide having RNA at its 3' end, is then ligated to the phosphate present at the 5' end of the decapped mRNA using T4 RNA ligase. The oligonucleotide may include a restriction site to facilitate cloning of the cDNAs following

their synthesis. Example 12 below describes one enzymatic method based on the doctoral thesis of Dumas.

## EXAMPLE 12

### Enzymatic Approach for Obtaining 5' ESTs

5 Twenty micrograms of PolyA+ RNA were dephosphorylated using Calf Intestinal Phosphatase (Biolabs). After a phenol chloroform extraction, the cap structure of mRNA was hydrolysed using the Tobacco Acid Pyrophosphatase (purified as described by Shinshi et al., Biochemistry 15: 2185-2190, 1976) and a hemi 5'DNA/RNA-3' oligonucleotide having an unphosphorylated 5' end, a stretch of adenosine ribophosphate at the 3' end, and an EcoRI site near the 5' end was ligated to the 5'P ends of mRNA using the T4 RNA ligase (Biolabs). Oligonucleotides suitable for use in this procedure are preferably 30-50 bases in length. Oligonucleotides having an unphosphorylated 5' end may be synthesized by adding a fluorochrome at the 5' end. The inclusion of a stretch of adenosine ribophosphates at the 3' end of the oligonucleotide increases ligation efficiency.

15 It will be appreciated that the oligonucleotide may contain cloning sites other than EcoRI.

Following ligation of the oligonucleotide to the phosphate present at the 5' end of the decapped mRNA, first and second strand cDNA synthesis may be carried out using conventional methods or those specified in EP0 625,572 and Kato et al. Construction of a Human Full-Length cDNA Bank. *Gene* 150:243-250 (1994), and Dumas Milne Edwards, *supra*, the disclosures of which are incorporated herein by reference. The resulting cDNA may then be ligated into vectors such as those disclosed in Kato et al. Construction of a Human Full-Length cDNA Bank. *Gene* 150:243-250 (1994) or other nucleic acid vectors known to those skilled in the art using techniques such as those described in Sambrook et al., *Molecular Cloning: A Laboratory Manual* 2d Ed., Cold Spring Harbor Laboratory Press, 1989, the disclosure of which is incorporated herein by reference.

## II. Characterization of 5' ESTs

The above chemical and enzymatic approaches for enriching mRNAs having intact 5' ends were employed to obtain 5' ESTs. First, mRNAs were prepared as described in Example 13 below.

### EXAMPLE 13

#### Preparation of mRNA

Total human RNAs or PolyA<sup>+</sup> RNAs derived from 29 different tissues were respectively purchased from LABIMO and CLONTECH and used to generate 44 cDNA libraries as described below. The purchased RNA had been isolated from cells or tissues using acid guanidium thiocyanate-phenol-chloroform extraction (Chomczynski, P and Sacchi, N., **Analytical Biochemistry** 162:156-159, 1987). PolyA<sup>+</sup> RNA was isolated from total RNA (LABIMO) by two passes of oligodT chromatography, as described by Aviv and Leder (Aviv, H. and Leder, P., **Proc. Natl. Acad. Sci. USA** 69:1408-1412, 1972) in order to eliminate ribosomal RNA.

The quality and the integrity of the poly A<sup>+</sup> were checked. Northern blots hybridized with a globin probe were used to confirm that the mRNAs were not degraded. Contamination of the PolyA<sup>+</sup> mRNAs by ribosomal sequences was checked using RNAs blots and a probe derived from the sequence of the 28S RNA. Preparations of mRNAs with less than 5% of ribosomal RNAs were used in library construction. To avoid constructing libraries with RNAs contaminated by exogenous sequences (prokaryotic or fungal), the presence of bacterial 16S ribosomal sequences or of two highly expressed mRNAs was examined using PCR.

Following preparation of the mRNAs, the above described chemical and/or the enzymatic procedures for enriching mRNAs having intact 5' ends discussed above were employed to obtain 5' ESTs from various tissues. In both approaches an oligonucleotide tag was attached to the cap at the 5' ends of the mRNAs. The oligonucleotide tag had an EcoRI site therein to facilitate later cloning procedures.

Following attachment of the oligonucleotide tag to the mRNA by either the chemical or enzymatic methods, the integrity of the mRNA was examined by performing a Northern blot with 200-500ng of mRNA using a probe complementary to the oligonucleotidetag.

### EXAMPLE 14

#### cDNA Synthesis Using mRNA Templates Having Intact 5' Ends

For the mRNAs joined to oligonucleotide tags using both the chemical and enzymatic methods, first strand cDNA synthesis was performed using reverse transcriptase with random nonamers as primers. In order to protect internal EcoRI sites in the cDNA from digestion at later steps in the procedure, methylated dCTP was used for first strand synthesis. After removal of RNA by an alkaline hydrolysis, the first strand of cDNA was precipitated using isopropanol in order to eliminate residual primers.

For both the chemical and the enzymatic methods, the second strand of the cDNA was synthesized with a Klenow fragment using a primer corresponding to the 5' end of the ligated oligonucleotide described in Example 12. Preferably, the primer is 20-25 bases in length. Methylated dCTP was also used for second strand synthesis in order to protect internal EcoRI sites in the cDNA from digestion during the cloning process.

Following cDNA synthesis, the cDNAs were cloned into pBlueScript as described in Example 15 below.

#### EXAMPLE 15

##### Insertion of cDNAs into BlueScript

Following second strand synthesis, the ends of the cDNA were blunted with T4 DNA polymerase (Biolabs) and the cDNA was digested with EcoRI. Since methylated dCTP was used during cDNA synthesis, the EcoRI site present in the tag was the only site which was hemi-methylated. Consequently, only the EcoRI site in the oligonucleotide tag was susceptible to EcoRI digestion. The cDNA was then size fractionated using exclusion chromatography (AcA, Biosepra). Fractions corresponding to cDNAs of more than 150 bp were pooled and ethanol precipitated. The cDNA was directionally cloned into the SmaI and EcoRI ends of the phagemid pBlueScript vector (Stratagene). The ligation mixture was electroporated into bacteria and propagated under appropriate antibiotic selection.

Clones containing the oligonucleotide tag attached were selected as described in Example 16 below.

#### EXAMPLE 16

##### Selection of Clones Having the Oligonucleotide Tag Attached Thereto

The plasmid DNAs containing 5' EST libraries made as described above were purified (Qiagen). A positive selection of the tagged clones was performed as follows. Briefly, in this selection procedure, the plasmid DNA was converted to single stranded DNA using gene II endonuclease of the phage F1 in combination with an exonuclease (Chang et al., **Gene** 127:95-8, 1993) such as exonuclease III or T7 gene 6 exonuclease. The resulting single stranded DNA was then purified using paramagnetic beads as described by Fry et al., **Biotechniques**, 13: 124-131, 1992. In this procedure, the single stranded DNA was hybridized with a biotinylated oligonucleotide having a sequence corresponding to the 3' end of the oligonucleotide described in Example 13. Preferably, the primer has a length of 20-25 bases. Clones including a sequence complementary to the biotinylated oligonucleotide were captured by incubation with streptavidin coated magnetic beads followed by magnetic selection. After capture of the positive clones, the plasmid DNA was released from the magnetic beads and converted into double stranded DNA using a DNA polymerase such as the ThermoSequenase obtained from Amersham Pharmacia Biotech. Alternatively, protocols such as the Gene Trapper kit (Gibco BRL) may be used. The double stranded DNA was then electroporated into bacteria. The percentage of positive clones having the 5' tag oligonucleotide was estimated to typically rank between 90 and 98% using dot blot analysis.

Following electroporation, the libraries were ordered in 384-microtiter plates (MTP). A copy of the MTP was stored for future needs. Then the libraries were transferred into 96 MTP and sequenced as described below.

### EXAMPLE 17

#### Sequencing of Inserts in Selected Clones

Plasmid inserts were first amplified by PCR on PE 9600 thermocyclers (Perkin-Elmer), using standard SETA-A and SETA-B primers (Genset SA), AmpliTaqGold (Perkin-Elmer), dNTPs (Boehringer), buffer and cycling conditions as recommended by the Perkin-Elmer Corporation.

PCR products were then sequenced using automatic ABI Prism 377 sequencers (Perkin Elmer, Applied Biosystems Division, Foster City, CA). Sequencing reactions

were performed using PE 9600 thermocyclers (Perkin Elmer) with standard dye-primer chemistry and ThermoSequenase (Amersham Life Science). The primers used were either T7 or 21M13 (available from Genset SA) as appropriate. The primers were labeled with the JOE, FAM, ROX and TAMRA dyes. The dNTPs and ddNTPs used in the sequencing reactions were purchased from Boehringer. Sequencing buffer, reagent concentrations and cycling conditions were as recommended by Amersham.

Following the sequencing reaction, the samples were precipitated with EtOH, resuspended in formamide loading buffer, and loaded on a standard 4% acrylamide gel. Electrophoresis was performed for 2.5 hours at 3000V on an ABI 377 sequencer, and the sequence data were collected and analyzed using the ABI Prism DNA Sequencing Analysis Software, version 2.1.2.

The sequence data from the 44 cDNA libraries made as described above were transferred to a proprietary database, where quality control and validation steps were performed. A proprietary base-caller ("Trace"), working using a Unix system automatically flagged suspect peaks, taking into account the shape of the peaks, the inter-peak resolution, and the noise level. The proprietary base-caller also performed an automatic trimming. Any stretch of 25 or fewer bases having more than 4 suspect peaks was considered unreliable and was discarded. Sequences corresponding to cloning vector or ligation oligonucleotides were automatically removed from the EST sequences. However, the resulting EST sequences may contain 1 to 5 bases belonging to the above mentioned sequences at their 5' end. If needed, these can easily be removed on a case by case basis.

Thereafter, the sequences were transferred to the proprietary NETGENE™ Database for further analysis as described below.

Following sequencing as described above, the sequences of the 5' ESTs were entered in a proprietary database called NETGENE™ for storage and manipulation. It will be appreciated by those skilled in the art that the data could be stored and manipulated on any medium which can be read and accessed by a computer. Computer readable media include magnetically readable media, optically readable media, or electronically readable

media. For example, the computer readable media may be a hard disc, a floppy disc, a magnetic tape, CD-ROM, RAM, or ROM as well as other types of other media known to those skilled in the art.

5 In addition, the sequence data may be stored and manipulated in a variety of data processor programs in a variety of formats. For example, the sequence data may be stored as text in a word processing file, such as MicrosoftWORD or WORDPERFECT or as an ASCII file in a variety of database programs familiar to those of skill in the art, such as DB2, SYBASE, or ORACLE.

10 The computer readable media on which the sequence information is stored may be in a personal computer, a network, a server or other computer systems known to those skilled in the art. The computer or other system preferably includes the storage media described above, and a processor for accessing and manipulating the sequence data.

15 Once the sequence data has been stored it may be manipulated and searched to locate those stored sequences which contain a desired nucleic acid sequence or which encode a protein having a particular functional domain. For example, the stored sequence information may be compared to other known sequences to identify homologies, motifs implicated in biological function, or structural motifs.

20 Programs which may be used to search or compare the stored sequences include the MacPattern (EMBL), BLAST, and BLAST2 program series (NCBI), basic local alignment search tool programs for nucleotide (BLASTN) and peptide (BLASTX) comparisons (Altschul et al, **J. Mol. Biol.** **215**: 403 (1990)) and FASTA (Pearson and Lipman, **Proc. Natl. Acad. Sci. USA**, **85**: 2444 (1988)). The BLAST programs then extend the alignments on the basis of defined match and mismatch criteria.

25 Motifs which may be detected using the above programs include sequences encoding leucine zippers, helix-turn-helix motifs, glycosylation sites, ubiquitination sites, alpha helices, and beta sheets, signal sequences encoding signal peptides which direct the secretion of the encoded proteins, sequences implicated in transcription regulation such as homeoboxes, acidic stretches, enzymatic active sites, substrate binding sites, and enzymatic cleavage sites.

Before searching the cDNAs in the NETGENE™ database for sequence motifs of interest, cDNAs derived from mRNAs which were not of interest were identified and eliminated from further consideration as described in Example 18 below.

### EXAMPLE 18

#### 5                    Elimination of Undesired Sequences from Further Consideration

5' ESTs in the NETGENE™ database which were derived from undesired sequences such as transfer RNAs, ribosomal RNAs, mitochondrial RNAs, procaryotic RNAs, fungal RNAs, Alu sequences, L1 sequences, or repeat sequences were identified using the FASTA and BLASTN programs with the parameters listed in Table II.

10                  To eliminate 5' ESTs encoding tRNAs from further consideration, the 5' EST sequences were compared to the sequences of 1190 known tRNAs obtained from EMBL release 38, of which 100 were human. The comparison was performed using FASTA on both strands of the 5' ESTs. Sequences having more than 80% homology over more than 60 nucleotides were identified as tRNA. Of the 144,341 sequences screened, 26 were  
15                  identified as tRNAs and eliminated from further consideration.

To eliminate 5' ESTs encoding rRNAs from further consideration, the 5' EST sequences were compared to the sequences of 2497 known rRNAs obtained from EMBL release 38, of which 73 were human. The comparison was performed using BLASTN on both strands of the 5' ESTs with the parameter S=108. Sequences having more than 80%  
20                  homology over stretches longer than 40 nucleotides were identified as rRNAs. Of the 144,341 sequences screened, 3,312 were identified as rRNAs and eliminated from further consideration.

To eliminate 5' ESTs encoding mtRNAs from further consideration, the 5' EST sequences were compared to the sequences of the two known mitochondrial genomes for which the entire genomic sequences are available and all sequences transcribed from these  
25                  mitochondrial genomes including tRNAs, rRNAs, and mRNAs for a total of 38 sequences. The comparison was performed using BLASTN on both strands of the 5' ESTs with the parameter S=108. Sequences having more than 80% homology over stretches longer than 40 nucleotides were identified as mtRNAs. Of the 144,341



sequences screened, 6,110 were identified as mtRNAs and eliminated from further consideration.

Sequences which might have resulted from exogenous contaminants were eliminated from further consideration by comparing the 5' EST sequences to release 46 of the EMBL bacterial and fungal divisions using BLASTN with the parameter S=144. All sequences having more than 90% homology over at least 40 nucleotides were identified as exogenous contaminants. Of the 42 cDNA libraries examined, the average percentages of procaryotic and fungal sequences contained therein were 0.2% and 0.5% respectively. Among these sequences, only one could be identified as a sequence specific to fungi. The others were either fungal or procaryotic sequences having homologies with vertebrate sequences or including repeat sequences which had not been masked during the electronic comparison.

In addition, the 5' ESTs were compared to 6093 Alu sequences and 1115 L1 sequences to mask 5' ESTs containing such repeat sequences from further consideration. 5' ESTs including THE and MER repeats, SSTR sequences or satellite, micro-satellite, or telomeric repeats were also eliminated from further consideration. On average, 11.5% of the sequences in the libraries contained repeat sequences. Of this 11.5%, 7% contained Alu repeats, 3.3% contained L1 repeats and the remaining 1.2% were derived from the other types of repetitive sequences which were screened. These percentages are consistent with those found in cDNA libraries prepared by other groups. For example, the cDNA libraries of Adams et al. contained between 0% and 7.4% Alu repeats depending on the source of the RNA which was used to prepare the cDNA library (Adams et al., *Nature* 377:174, 1996).

The sequences of those 5' ESTs remaining after the elimination of undesirable sequences were compared with the sequences of known human mRNAs to determine the accuracy of the sequencing procedures described above.

#### EXAMPLE 19

##### Measurement of Sequencing Accuracy by Comparison to Known Sequences

To further determine the accuracy of the sequencing procedure described above, the sequences of 5' ESTs derived from known sequences were identified and compared to the known sequences. First, a FASTA analysis with overhangs shorter than 5 bp on both ends was conducted on the 5' ESTs to identify those matching an entry in the public human mRNA database. The 6655 5' ESTs which matched a known human mRNA were then realigned with their cognate mRNA and dynamic programming was used to include substitutions, insertions, and deletions in the list of "errors" which would be recognized. Errors occurring in the last 10 bases of the 5' EST sequences were ignored to avoid the inclusion of spurious cloning sites in the analysis of sequencing accuracy.

This analysis revealed that the sequences incorporated in the NETGENE™ database had an accuracy of more than 99.5%.

To determine the efficiency with which the above selection procedures select cDNAs which include the 5' ends of their corresponding mRNAs, the following analysis was performed.

#### EXAMPLE 20

##### Determination of Efficiency of 5' EST Selection

To determine the efficiency at which the above selection procedures isolated 5' ESTs which included sequences close to the 5' end of the mRNAs from which they were derived, the sequences of the ends of the 5' ESTs which were derived from the elongation factor 1 subunit  $\alpha$  and ferritin heavy chain genes were compared to the known cDNA sequences for these genes. Since the transcription start sites for the elongation factor 1 subunit  $\alpha$  and ferritin heavy chain are well characterized, they may be used to determine the percentage of 5' ESTs derived from these genes which included the authentic transcription start sites.

For both genes, more than 95% of the cDNAs included sequences close to or upstream of the 5' end of the corresponding mRNAs.

To extend the analysis of the reliability of the procedures for isolating 5' ESTs from ESTs in the NETGENE™ database, a similar analysis was conducted using a database composed of human mRNA sequences extracted from GenBank database release

97 for comparison. For those 5' ESTs derived from mRNAs included in the GeneBank database, more than 85% had their 5' ends close to the 5' ends of the known sequence. As some of the mRNA sequences available in the GenBank database are deduced from genomic sequences, a 5' end matching with these sequences will be counted as an internal match. Thus, the method used here underestimates the yield of ESTs including the authentic 5' ends of their corresponding mRNAs.

The EST libraries made above included multiple 5' ESTs derived from the same mRNA. The sequences of such 5' ESTs were compared to one another and the longest 5' ESTs for each mRNA were identified. Overlapping cDNAs were assembled into continuous sequences (contigs). The resulting continuous sequences were then compared to public databases to gauge their similarity to known sequences, as described in Example 21 below.

### EXAMPLE 21

#### Clustering of the 5' ESTs and Calculation of Novelty Indices for cDNA Libraries

For each sequenced EST library, the sequences were clustered by the 5' end. Each sequence in the library was compared to the others with BLASTN2 (direct strand, parameters S=107). ESTs with High Scoring Segment Pairs (HSPs) at least 25 bp long, having 95% identical bases and beginning closer than 10 bp from each EST 5' end were grouped. The longest sequence found in the cluster was used as representative of the cluster. A global clustering between libraries was then performed leading to the definition of super-contigs.

To assess the yield of new sequences within the EST libraries, a novelty rate (NR) was defined as:  $NR = 100 \times (\text{Number of new unique sequences found in the library} / \text{Total number of sequences from the library})$ . Typically, novelty rating range between 10% and 41% depending on the tissue from which the EST library was obtained. For most of the libraries, the random sequencing of 5' EST libraries was pursued until the novelty rate reached 20%.

Following characterization as described above, the collection of 5' ESTs in NETGENE™ was screened to identify those 5' ESTs bearing potential signal sequences as described in Example 22 below.

## EXAMPLE 22

### 5                    Identification of Potential Signal Sequences in 5' ESTs

10                    The 5' ESTs in the NETGENE™ database were screened to identify those having an uninterrupted open reading frame (ORF) longer than 45 nucleotides beginning with an ATG codon and extending to the end of the EST. Approximately half of the cDNA sequences in NETGENE™ contained such an ORF. The ORFs of these 5' ESTs were  
15                    searched to identify potential signal motifs using slight modifications of the procedures disclosed in Von Heijne, G. A New Method for Predicting Signal Sequence Cleavage Sites. **Nucleic Acids Res.** 14:4683-4690 (1986), the disclosure of which is incorporated herein by reference. Those 5' EST sequences encoding a 15 amino acid long stretch with a score of at least 3.5 in the Von Heijne signal peptide identification matrix were  
20                    considered to possess a signal sequence. Those 5' ESTs which matched a known human mRNA or EST sequence and had a 5' end more than 20 nucleotides downstream of the known 5' end were excluded from further analysis. The remaining cDNAs having signal sequences therein were included in a database called SIGNALTAG™.

20                    To confirm the accuracy of the above method for identifying signal sequences, the analysis of Example 23 was performed.

## EXAMPLE 23

### Confirmation of Accuracy of Identification of Potential Signal Sequences in 5' ESTs

25                    The accuracy of the above procedure for identifying signal sequences encoding signal peptides was evaluated by applying the method to the 43 amino terminal amino acids of all human SwissProt proteins. The computed Von Heijne score for each protein was compared with the known characterization of the protein as being a secreted protein or a non-secreted protein. In this manner, the number of non-secreted proteins having a score higher than 3.5 (false positives) and the number of secreted proteins having a score lower than 3.5 (false negatives) could be calculated.

Using the results of the above analysis, the probability that a peptide encoded by the 5' region of the mRNA is in fact a genuine signal peptide based on its Von Heijne's score was calculated based on either the assumption that 10% of human proteins are secreted or the assumption that 20% of human proteins are secreted. The results of this analysis are shown in Figures 2 and 3.

Using the above method of identifying secretory proteins, 5' ESTs for human glucagon, gamma interferon induced monokine precursor, secreted cyclophilin-like protein, human pleiotropin, and human biotinidase precursor all of which are polypeptides which are known to be secreted, were obtained. Thus, the above method successfully identified those 5' ESTs which encode a signal peptide.

To confirm that the signal peptide encoded by the 5' ESTs actually functions as a signal peptide, the signal sequences from the 5' ESTs may be cloned into a vector designed for the identification of signal peptides. Some signal peptide identification vectors are designed to confer the ability to grow in selective medium on host cells which have a signal sequence operably inserted into the vector. For example, to confirm that a 5' EST encodes a genuine signal peptide, the signal sequence of the 5' EST may be inserted upstream and in frame with a non-secreted form of the yeast invertase gene in signal peptide selection vectors such as those described in U.S. Patent No. 5,536,637, the disclosure of which is incorporated herein by reference. Growth of host cells containing signal sequence selection vectors having the signal sequence from the 5' EST inserted therein confirms that the 5' EST encodes a genuine signal peptide.

Alternatively, the presence of a signal peptide may be confirmed by cloning the extended cDNAs obtained using the ESTs into expression vectors such as pXT1 (as described below), or by constructing promoter-signal sequence-reporter gene vectors which encode fusion proteins between the signal peptide and an assayable reporter protein. After introduction of these vectors into a suitable host cell, such as COS cells or NIH 3T3 cells, the growth medium may be harvested and analyzed for the presence of the secreted protein. The medium from these cells is compared to the medium from cells containing

vectors lacking the signal sequence or extended cDNA insert to identify vectors which encode a functional signal peptide or an authentic secreted protein.

Those 5' ESTs which encoded a signal peptide, as determined by the method of Example 22 above, were further grouped into four categories based on their homology to known sequences. The categorization of the 5' ESTs is described in Example 24 below.

#### EXAMPLE 24

##### Categorization of 5' ESTs Encoding a Signal Peptide

Those 5' ESTs having a sequence not matching any known vertebrate sequence nor any publicly available EST sequence were designated "new." Of the sequences in the SIGNALTAG™ database, 947 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those 5' ESTs having a sequence not matching any vertebrate sequence but matching a publicly known EST were designated "EST-ext", provided that the known EST sequence was extended by at least 40 nucleotides in the 5' direction. Of the sequences in the SIGNALTAG™ database, 150 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those ESTs not matching any vertebrate sequence but matching a publicly known EST without extending the known EST by at least 40 nucleotides in the 5' direction were designated "EST." Of the sequences in the SIGNALTAG™ database, 599 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category.

Those 5' ESTs matching a human mRNA sequence but extending the known sequence by at least 40 nucleotides in the 5' direction were designated "VERT-ext." Of the sequences in the SIGNALTAG™ database, 23 of the 5' ESTs having a Von Heijne's score of at least 3.5 fell into this category. Included in this category was a 5' EST which extended the known sequence of the human translocase mRNA by more than 200 bases in the 5' direction. A 5' EST which extended the sequence of a human tumor suppressor gene in the 5' direction was also identified.

Figure 4 shows the distribution of 5' ESTs in each category and the number of 5' ESTs in each category having a given minimum von Heijne's score.

Each of the 5' ESTs was categorized based on the tissue from which its corresponding mRNA was obtained, as described below in Example 25.

### EXAMPLE 25

#### Categorization of Expression Patterns

5        Figure 5 shows the tissues from which the mRNAs corresponding to the 5' ESTs in each of the above described categories were obtained.

      In addition to categorizing the 5' ESTs by the tissue from which the cDNA library in which they were first identified was obtained, the spatial and temporal expression patterns of the mRNAs corresponding to the 5' ESTs, as well as their expression levels, 10        may be determined as described in Example 26 below. Characterization of the spatial and temporal expression patterns and expression levels of these mRNAs is useful for constructing expression vectors capable of producing a desired level of gene product in a desired spatial or temporal manner, as will be discussed in more detail below.

      In addition, 5' ESTs whose corresponding mRNAs are associated with disease 15        states may also be identified. For example, a particular disease may result from lack of expression, over expression, or under expression of an mRNA corresponding to a 5' EST. By comparing mRNA expression patterns and quantities in samples taken from healthy individuals with those from individuals suffering from a particular disease, 5' ESTs responsible for the disease may be identified.

20        It will be appreciated that the results of the above characterization procedures for 5' ESTs also apply to extended cDNAs (obtainable as described below) which contain sequences adjacent to the 5' ESTs. It will also be appreciated that if it is desired to defer characterization until extended cDNAs have been obtained rather than characterizing the ESTs themselves, the above characterization procedures can be applied to characterize the 25        extended cDNAs after their isolation.

### EXAMPLE 26

#### Evaluation of Expression Levels and Patterns of mRNAs

#### Corresponding to 5' ESTs or Extended cDNAs

Expression levels and patterns of mRNAs corresponding to 5' ESTs or extended cDNAs (obtainable as described below) may be analyzed by solution hybridization with long probes as described in International Patent Application No. WO 97/05277, the entire contents of which are hereby incorporated by reference. Briefly, a 5' EST, extended  
5 cDNA, or fragment thereof corresponding to the gene encoding the mRNA to be characterized is inserted at a cloning site immediately downstream of a bacteriophage (T3, T7 or SP6) RNA polymerase promoter to produce antisense RNA. Preferably, the 5' EST or extended cDNA has 100 or more nucleotides. The plasmid is linearized and transcribed in the presence of ribonucleotides comprising modified ribonucleotides (i.e. biotin-UTP and DIG-UTP). An excess of this doubly labeled RNA is hybridized in solution with  
10 mRNA isolated from cells or tissues of interest. The hybridizations are performed under standard stringent conditions (40-50°C for 16 hours in an 80% formamide, 0.4 M NaCl buffer, pH 7-8). The unhybridized probe is removed by digestion with ribonucleases specific for single-stranded RNA (i.e. RNases CL3, T1, Phy M, U2 or A). The presence  
15 of the biotin-UTP modification enables capture of the hybrid on a microtitration plate coated with streptavidin. The presence of the DIG modification enables the hybrid to be detected and quantified by ELISA using an anti-DIG antibody coupled to alkaline phosphatase.

The 5' ESTs, extended cDNAs, or fragments thereof may also be tagged with  
20 nucleotide sequences for the serial analysis of gene expression (SAGE) as disclosed in UK Patent Application No. 2 305 241 A, the entire contents of which are incorporated by reference. In this method, cDNAs are prepared from a cell, tissue, organism or other source of nucleic acid for which it is desired to determine gene expression patterns. The resulting cDNAs are separated into two pools. The cDNAs in each pool are cleaved with  
25 a first restriction endonuclease, called an "anchoring enzyme," having a recognition site which is likely to be present at least once in most cDNAs. The fragments which contain the 5' or 3' most region of the cleaved cDNA are isolated by binding to a capture medium such as streptavidin coated beads. A first oligonucleotide linker having a first sequence for hybridization of an amplification primer and an internal restriction site for a "tagging



endonuclease" is ligated to the digested cDNAs in the first pool. Digestion with the second endonuclease produces short "tag" fragments from the cDNAs.

A second oligonucleotide having a second sequence for hybridization of an amplification primer and an internal restriction site is ligated to the digested cDNAs in the second pool. The cDNA fragments in the second pool are also digested with the "tagging endonuclease" to generate short "tag" fragments derived from the cDNAs in the second pool. The "tags" resulting from digestion of the first and second pools with the anchoring enzyme and the tagging endonuclease are ligated to one another to produce "ditags." In some embodiments, the ditags are concatamerized to produce ligation products containing from 2 to 200 ditags. The tag sequences are then determined and compared to the sequences of the 5' ESTs or extended cDNAs to determine which 5' ESTs or extended cDNAs are expressed in the cell, tissue, organism, or other source of nucleic acids from which the tags were derived. In this way, the expression pattern of the 5' ESTs or extended cDNAs in the cell, tissue, organism, or other source of nucleic acids is obtained.

Quantitative analysis of gene expression may also be performed using arrays. As used herein, the term array means a one dimensional, two dimensional, or multidimensional arrangement of full length cDNAs (i.e. extended cDNAs which include the coding sequence for the signal peptide, the coding sequence for the mature protein, and a stop codon), extended cDNAs, 5' ESTs or fragments of the full length cDNAs, extended cDNAs, or 5' ESTs of sufficient length to permit specific detection of gene expression. Preferably, the fragments are at least 15 nucleotides in length. More preferably, the fragments are at least 100 nucleotides in length. More preferably, the fragments are more than 100 nucleotides in length. In some embodiments the fragments may be more than 500 nucleotides in length.

For example, quantitative analysis of gene expression may be performed with full length cDNAs, extended cDNAs, 5' ESTs, or fragments thereof in a complementary DNA microarray as described by Schena et al. (*Science* **270**:467-470, 1995; *Proc. Natl. Acad. Sci. U.S.A.* **93**:10614-10619, 1996). Full length cDNAs, extended cDNAs, 5' ESTs or fragments thereof are amplified by PCR and arrayed from 96-well microtiter plates onto

silylated microscope slides using high-speed robotics. Printed arrays are incubated in a humid chamber to allow rehydration of the array elements and rinsed, once in 0.2% SDS for 1 min, twice in water for 1 min and once for 5 min in sodium borohydride solution. The arrays are submerged in water for 2 min at 95°C, transferred into 0.2% SDS for 1 min, rinsed twice with water, air dried and stored in the dark at 25°C.

Cell or tissue mRNA is isolated or commercially obtained and probes are prepared by a single round of reverse transcription. Probes are hybridized to 1 cm<sup>2</sup> microarrays under a 14 x 14 mm glass coverslip for 6-12 hours at 60°C. Arrays are washed for 5 min at 25°C in low stringency wash buffer (1 x SSC/0.2% SDS), then for 10 min at room temperature in high stringency wash buffer (0.1 x SSC/0.2% SDS). Arrays are scanned in 0.1 x SSC using a fluorescence laser scanning device fitted with a custom filter set. Accurate differential expression measurements are obtained by taking the average of the ratios of two independent hybridizations.

Quantitative analysis of the expression of genes may also be performed with full length cDNAs, extended cDNAs, 5' ESTs, or fragments thereof in complementary DNA arrays as described by Pietu et al. (Genome Research 6:492-503, 1996). The full length cDNAs, extended cDNAs, 5' ESTs or fragments thereof are PCR amplified and spotted on membranes. Then, mRNAs originating from various tissues or cells are labeled with radioactive nucleotides. After hybridization and washing in controlled conditions, the hybridized mRNAs are detected by phospho-imaging or autoradiography. Duplicate experiments are performed and a quantitative analysis of differentially expressed mRNAs is then performed.

Alternatively, expression analysis of the 5' ESTs or extended cDNAs can be done through high density nucleotide arrays as described by Lockhart et al. (Nature Biotechnology 14: 1675-1680, 1996) and Sosnowsky et al. (Proc. Natl. Acad. Sci. 94:1119-1123, 1997). Oligonucleotides of 15-50 nucleotides corresponding to sequences of the 5' ESTs or extended cDNAs are synthesized directly on the chip (Lockhart et al., *supra*) or synthesized and then addressed to the chip (Sosnowski et al., *supra*). Preferably, the oligonucleotides are about 20 nucleotides in length.

cDNA probes labeled with an appropriate compound, such as biotin, digoxigenin or fluorescent dye, are synthesized from the appropriate mRNA population and then randomly fragmented to an average size of 50 to 100 nucleotides. The said probes are then hybridized to the chip. After washing as described in Lockhart et al., *supra* and application of different electric fields (Sosnowsky et al., Proc. Natl. Acad. Sci. 94:1119-1123), the dyes or labeling compounds are detected and quantified. Duplicate hybridizations are performed. Comparative analysis of the intensity of the signal originating from cDNA probes on the same target oligonucleotide in different cDNA samples indicates a differential expression of the mRNA corresponding to the 5' EST or extended cDNA from which the oligonucleotide sequence has been designed.

### **III. Use of 5' ESTs to Clone Extended cDNAs and to Clone the Corresponding Genomic DNAs**

Once 5' ESTs which include the 5' end of the corresponding mRNAs have been selected using the procedures described above, they can be utilized to isolate extended cDNAs which contain sequences adjacent to the 5' ESTs. The extended cDNAs may include the entire coding sequence of the protein encoded by the corresponding mRNA, including the authentic translation start site, the signal sequence, and the sequence encoding the mature protein remaining after cleavage of the signal peptide. Such extended cDNAs are referred to herein as "full length cDNAs." Alternatively, the extended cDNAs may include only the sequence encoding the mature protein remaining after cleavage of the signal peptide, or only the sequence encoding the signal peptide.

Example 27 below describes a general method for obtaining extended cDNAs. Example 28 below describes the cloning and sequencing of several extended cDNAs, including extended cDNAs which include the entire coding sequence and authentic 5' end of the corresponding mRNA for several secreted proteins.

The methods of Examples 27, 28, and 29 can also be used to obtain extended cDNAs which encode less than the entire coding sequence of the secreted proteins encoded by the genes corresponding to the 5' ESTs. In some embodiments, the extended cDNAs isolated using these methods encode at least 10 amino acids of one of the proteins

encoded by the sequences of SEQ ID NOs: 40-140 and 242-377. In further embodiments, the extended cDNAs encode at least 20 amino acids of the proteins encoded by the sequences of SEQ ID NOs: 40-140 and 242-377. In further embodiments, the extended cDNAs encode at least 30 amino acids of the sequences of SEQ ID NOs: 40-140 and 242-377. In a preferred embodiment, the extended cDNAs encode a full length protein sequence, which includes the protein coding sequences of SEQ ID NOs: 40-140 and 242-377.

## EXAMPLE 27

### General Method for Using 5' ESTs to Clone and Sequence Extended cDNAs

The following general method has been used to quickly and efficiently isolate extended cDNAs including sequence adjacent to the sequences of the 5' ESTs used to obtain them. This method may be applied to obtain extended cDNAs for any 5' EST in the NETGENE™ database, including those 5' ESTs encoding secreted proteins. The method is summarized in Figure 6.

#### 1. Obtaining Extended cDNAs

##### a) First strand synthesis

The method takes advantage of the known 5' sequence of the mRNA. A reverse transcription reaction is conducted on purified mRNA with a poly 14dT primer containing a 49 nucleotide sequence at its 5' end allowing the addition of a known sequence at the end of the cDNA which corresponds to the 3' end of the mRNA. For example, the primer may have the following sequence: 5'-ATC GTT GAG ACT CGT ACC AGC AGA GTC ACG AGA GAG ACT ACA CGG TAC TGG TTT TTT TTT TTT TTVN -3' (SEQ ID NO:14). Those skilled in the art will appreciate that other sequences may also be added to the poly dT sequence and used to prime the first strand synthesis. Using this primer and a reverse transcriptase such as the Superscript II (Gibco BRL) or Rnase H Minus M-MLV (Promega) enzyme, a reverse transcript anchored at the 3' polyA site of the RNAs is generated.

After removal of the mRNA hybridized to the first cDNA strand by alkaline hydrolysis, the products of the alkaline hydrolysis and the residual poly dT primer are

eliminated with an exclusion column such as an AcA34 (Biosepra) matrix as explained in Example 11.

b) Second strand synthesis

A pair of nested primers on each end is designed based on the known 5' sequence from the 5' EST and the known 3' end added by the poly dT primer used in the first strand synthesis. Software used to design primers are either based on GC content and melting temperatures of oligonucleotides, such as OSP (Illier and Green, *PCR Meth. Appl.* **1**:124-128, 1991), or based on the octamer frequency disparity method (Griffais et al., *Nucleic Acids Res.* **19**: 3887-3891, 1991 such as PC-Rare

(<http://bioinformatics.weizmann.ac.il/software/PC-Rare/doc/manuel.html>).

Preferably, the nested primers at the 5' end are separated from one another by four to nine bases. The 5' primer sequences may be selected to have melting temperatures and specificities suitable for use in PCR.

Preferably, the nested primers at the 3' end are separated from one another by four to nine bases. For example, the nested 3' primers may have the following sequences: (5'-CCA GCA GAG TCA CGA GAG AGA CTA CAC GG -3'(SEQ ID NO:15), and 5'-CAC GAG AGA GAC TAC ACG GTA CTG G -3' (SEQ ID NO:16). These primers were selected because they have melting temperatures and specificities compatible with their use in PCR. However, those skilled in the art will appreciate that other sequences may also be used as primers.

The first PCR run of 25 cycles is performed using the Advantage Tth Polymerase Mix (Clontech) and the outer primer from each of the nested pairs. A second 20 cycle PCR using the same enzyme and the inner primer from each of the nested pairs is then performed on 1/2500 of the first PCR product. Thereafter, the primers and nucleotides are removed.

2. Sequencing of Full Length Extended cDNAs or Fragments Thereof

Due to the lack of position constraints on the design of 5' nested primers compatible for PCR use using the OSP software, amplicons of two types are obtained. Preferably, the second 5' primer is located upstream of the translation initiation codon

thus yielding a nested PCR product containing the whole coding sequence. Such a full length extended cDNA undergoes a direct cloning procedure as described in section a below. However, in some cases, the second 5' primer is located downstream of the translation initiation codon, thereby yielding a PCR product containing only part of the ORF. Such incomplete PCR products are submitted to a modified procedure described in section b below.

a) Nested PCR products containing complete ORFs

When the resulting nested PCR product contains the complete coding sequence, as predicted from the 5' EST sequence, it is cloned in an appropriate vector such as pED6dpc2, as described in section 3.

b) Nested PCR products containing incomplete ORFs

When the amplicon does not contain the complete coding sequence, intermediate steps are necessary to obtain both the complete coding sequence and a PCR product containing the full coding sequence. The complete coding sequence can be assembled from several partial sequences determined directly from different PCR products as described in the following section.

Once the full coding sequence has been completely determined, new primers compatible for PCR use are designed to obtain amplicons containing the whole coding region. However, in such cases, 3' primers compatible for PCR use are located inside the 3' UTR of the corresponding mRNA, thus yielding amplicons which lack part of this region, i.e. the polyA tract and sometimes the polyadenylation signal, as illustrated in figure 6. Such full length extended cDNAs are then cloned into an appropriate vector as described in section 3.

c) Sequencing extended cDNAs

Sequencing of extended cDNAs is performed using a Die Terminator approach with the AmpliTaq DNA polymerase FS kit available from Perkin Elmer.

In order to sequence PCR fragments, primer walking is performed using software such as OSP to choose primers and automated computer software such as ASMG (Sutton et al., *Genome Science Technol.* 1: 9-19, 1995) to construct contigs of walking sequences

including the initial 5' tag using minimum overlaps of 32 nucleotides. Preferably, primer walking is performed until the sequences of full length cDNAs are obtained.

Completion of the sequencing of a given extended cDNA fragment is assessed as follows. Since sequences located after a polyA tract are difficult to determine precisely in the case of uncloned products, sequencing and primer walking processes for PCR products are interrupted when a polyA tract is identified in extended cDNAs obtained as described in case b. The sequence length is compared to the size of the nested PCR product obtained as described above. Due to the limited accuracy of the determination of the PCR product size by gel electrophoresis, a sequence is considered complete if the size of the obtained sequence is at least 70 % the size of the first nested PCR product. If the length of the sequence determined from the computer analysis is not at least 70% of the length of the nested PCR product, these PCR products are cloned and the sequence of the insertion is determined. When Northern blot data are available, the size of the mRNA detected for a given PCR product is used to finally assess that the sequence is complete. Sequences which do not fulfill the above criteria are discarded and will undergo a new isolation procedure.

Sequence data of all extended cDNAs are then transferred to a proprietary database, where quality controls and validation steps are carried out as described in example 15.

### 20 3. Cloning of Full Length Extended cDNAs

The PCR product containing the full coding sequence is then cloned in an appropriate vector. For example, the extended cDNAs can be cloned into the expression vector pED6dpc2 (DiscoverEase, Genetics Institute, Cambridge, MA) as follows. The structure of pED6dpc2 is shown in Figure 7. pED6dpc2 vector DNA is prepared with blunt ends by performing an EcoRI digestion followed by a fill in reaction. The blunt ended vector is dephosphorylated. After removal of PCR primers and ethanol precipitation, the PCR product containing the full coding sequence or the extended cDNA obtained as described above is phosphorylated with a kinase subsequently removed by phenol-Sevag extraction and precipitation. The double stranded extended cDNA is then

ligated to the vector and the resulting expression plasmid introduced into appropriate host cells.

Since the PCR products obtained as described above are blunt ended molecules that can be cloned in either direction, the orientation of several clones for each PCR product is determined. Then, 4 to 10 clones are ordered in microtiter plates and subjected to a PCR reaction using a first primer located in the vector close to the cloning site and a second primer located in the portion of the extended cDNA corresponding to the 3' end of the mRNA. This second primer may be the antisense primer used in anchored PCR in the case of direct cloning (case a) or the antisense primer located inside the 3'UTR in the case of indirect cloning (case b). Clones in which the start codon of the extended cDNA is operably linked to the promoter in the vector so as to permit expression of the protein encoded by the extended cDNA are conserved and sequenced. In addition to the ends of cDNA inserts, approximately 50 bp of vector DNA on each side of the cDNA insert are also sequenced.

The cloned PCR products are then entirely sequenced according to the aforementioned procedure. In this case, contig assembly of long fragments is then performed on walking sequences that have already contigated for uncloned PCR products during primer walking. Sequencing of cloned amplicons is complete when the resulting contigs include the whole coding region as well as overlapping sequences with vector DNA on both ends.

#### 4. Computer Analysis of Full Length Extended cDNA

Sequences of all full length extended cDNAs are then submitted to further analysis as described below and using the parameters found in Table II with the following modifications. For screening of miscellaneous subdivisions of Genbank, FASTA was used instead of BLASTN and 15 nucleotide of homology was the limit instead of 17. For Alu detection, BLASTN was used with the following parameters: S=72; identity=70%; and length = 40 nucleotides. Polyadenylation signal and polyA tail which were not search for the 5' ESTs were searched. For polyadenylation signal detection the signal (AATAAA) was searched with one permissible mismatch in the last



ten nucleotides preceding the 5' end of the polyA. For the polyA, a stretch of 8 amino acids in the last 20 nucleotides of the sequence was searched with BLAST2N in the sense strand with the following parameters (W=6, S=10, E=1000, and identity=90%). Finally, patented sequences and ORF homologies were searched using, respectively,

5 BLASTN and BLASTP on GenSEQ (Derwent's database of patented nucleotide sequences) and SWISSPROT for ORFs with the following parameters (W=8 and B=10). Before examining the extended full length cDNAs for sequences of interest, extended cDNAs which are not of interest are searched as follows.

a) Elimination of undesired sequences

10 Although 5'ESTs were checked to remove contaminants sequences as described in Example 18, a last verification was carried out to identify extended cDNAs sequences derived from undesired sequences such as vector RNAs, transfer RNAs, ribosomal rRNAs, mitochondrial RNAs, prokaryotic RNAs and fungal RNAs using the FASTA and BLASTN programs on both strands of extended cDNAs as described below.

15 To identify the extended cDNAs encoding vector RNAs, extended cDNAs are compared to the known sequences of vector RNA using the FASTA program. Sequences of extended cDNAs with more than 90% homology over stretches of 15 nucleotides are identified as vector RNA.

20 To identify the extended cDNAs encoding tRNAs, extended cDNA sequences were compared to the sequences of 1190 known tRNAs obtained from EMBL release 38, of which 100 were human. Sequences of extended cDNAs having more than 80% homology over 60 nucleotides using FASTA were identified as tRNA.

25 To identify the extended cDNAs encoding rRNAs, extended cDNA sequences were compared to the sequences of 2497 known rRNAs obtained from EMBL release 38, of which 73 were human. Sequences of extended cDNAs having more than 80% homology over stretches longer than 40 nucleotides using BLASTN were identified as rRNAs.

To identify the extended cDNAs encoding mtRNAs, extended cDNA sequences were compared to the sequences of the two known mitochondrial genomes for which the

entire genomic sequences are available and all sequences transcribed from these mitochondrial genomes including tRNAs, rRNAs, and mRNAs for a total of 38 sequences. Sequences of extended cDNAs having more than 80% homology over stretches longer than 40 nucleotides using BLASTN were identified as mtRNAs.

5 Sequences which might have resulted from other exogenous contaminants were identified by comparing extended cDNA sequences to release 105 of Genbank bacterial and fungal divisions. Sequences of extended cDNAs having more than 90% homology over 40 nucleotides using BLASTN were identified as exogenous prokaryotic or fungal contaminants.

10 In addition, extended cDNAs were searched for different repeat sequences, including Alu sequences, L1 sequences, THE and MER repeats, SSTR sequences or satellite, micro-satellite, or telomeric repeats. Sequences of extended cDNAs with more than 70% homology over 40 nucleotide stretches using BLASTN were identified as repeat sequences and masked in further identification procedures. In addition, clones  
15 showing extensive homology to repeats, i.e., matches of either more than 50 nucleotides if the homology was at least 75% or more than 40 nucleotides if the homology was at least 85% or more than 30 nucleotides if the homology was at least 90%, were flagged.

#### b) Identification of structural features

20 Structural features, e.g. polyA tail and polyadenylation signal, of the sequences of full length extended cDNAs are subsequently determined as follows.

A polyA tail is defined as a homopolymeric stretch of at least 11 A with at most one alternative base within it. The polyA tail search is restricted to the last 100 nt of the sequence and limited to stretches of 11 consecutive A's because sequencing reactions  
25 are often not readable after such a polyA stretch. Stretches having more than 90% homology over 8 nucleotides are identified as polyA tails using BLAST2N.

A polyA tail is defined as a homopolymeric stretch of at least 11 A with at most one alternative base within it. The polyA tail search is restricted to the last 20 nt of the sequence and limited to stretches of 11 consecutive A's because sequencing reactions

are often not readable after such a polyA stretch. Stretches with 100% homology over 6 nucleotides are identified as polyA tails.

To search for a polyadenylation signal, the polyA tail is clipped from the full-length sequence. The 50 bp preceding the polyA tail are searched for the canonic polyadenylation AAUAAA signal allowing one mismatch to account for possible sequencing errors and known variation in the canonical sequence of the polyadenylation signal.

#### c) Identification of functional features

Functional features, e.g. ORFs and signal sequences, of the sequences of full length extended cDNAs were subsequently determined as follows.

The 3 upper strand frames of extended cDNAs are searched for ORFs defined as the maximum length fragments beginning with a translation initiation codon and ending with a stop codon. ORFs encoding at least 20 amino acids are preferred.

Each found ORF is then scanned for the presence of a signal peptide in the first 50 amino-acids or, where appropriate, within shorter regions down to 20 amino acids or less in the ORF, using the matrix method of von Heijne (Nuc. Acids Res. 14: 4683-4690 (1986)), the disclosure of which is incorporated herein by reference and the modification described in Example 22.

#### d) Homology to either nucleotidic or proteic sequences

Sequences of full length extended cDNAs are then compared to known sequences on a nucleotidic or proteic basis.

Sequences of full length extended cDNAs are compared to the following known nucleic acid sequences: vertebrate sequences, EST sequences, patented sequences and recently identified sequences available at the time of filing the priority documents. Full length cDNA sequences are also compared to the sequences of a private database (Genset internal sequences) in order to find sequences that have already been identified by applicants. Sequences of full length extended cDNAs with more than 90% homology over 30 nucleotides using either BLASTN or BLAST2N as indicated in Table III are identified as sequences that have already been described. Matching vertebrate sequences

are subsequently examined using FASTA; full length extended cDNAs with more than 70% homology over 30 nucleotides are identified as sequences that have already been described.

5 ORFs encoded by full length extended cDNAs as defined in section c) are subsequently compared to known amino acid sequences found in Swissprot release CHP, PIR release PIR# and Genpept release GPEPT public databases using BLASTP with the parameter W=8 and allowing a maximum of 10 matches. Sequences of full length extended cDNAs showing extensive homology to known protein sequences are recognized as already identified proteins.

10 In addition, the three-frame conceptual translation products of the top strand of full length extended cDNAs are compared to publicly known amino acid sequences of Swissprot using BLASTX with the parameter E=0.001. Sequences of full length extended cDNAs with more than 70% homology over 30 amino acid stretches are detected as already identified proteins.

15 5. Selection of Cloned Full Length Sequences of the Present Invention

Cloned full length extended cDNA sequences that have already been characterized by the aforementioned computer analysis are then submitted to an automatic procedure in order to preselect full length extended cDNAs containing sequences of interest.

20 a) Automatic sequence preselection

All complete cloned full length extended cDNAs clipped for vector on both ends are considered. First, a negative selection is operated in order to eliminate unwanted cloned sequences resulting from either contaminants or PCR artifacts as follows. Sequences matching contaminant sequences such as vector RNA, tRNA, mtRNA, rRNA  
25 sequences are discarded as well as those encoding ORF sequences exhibiting extensive homology to repeats as defined in section 4 a). Sequences obtained by direct cloning using nested primers on 5' and 3' tags (section 1. case a) but lacking polyA tail are discarded. Only ORFs containing a signal peptide and ending either before the polyA tail (case a) or before the end of the cloned 3'UTR (case b) are kept. Then, ORFs containing

unlikely mature proteins such as mature proteins which size is less than 20 amino acids or less than 25% of the immature protein size are eliminated.

In the selection of the OFR, priority was given to the ORF and the frame corresponding to the polypeptides described in SignalTag Patents (United States Patent Application Serial Nos: 08/905,223; 08/905,135; 08/905,051; 08/905,144; 08/905,279; 08/904,468; 08/905,134; and 08/905,133). If the ORF was not found among the OFRs described in the SignalTag Patents, the ORF encoding the signal peptide with the highest score according to Von Heijne method as defined in Example 22 was chosen. If the scores were identical, then the longest ORF was chosen.

Sequences of full length extended cDNA clones are then compared pairwise with BLAST after masking of the repeat sequences. Sequences containing at least 90% homology over 30 nucleotides are clustered in the same class. Each cluster is then subjected to a cluster analysis that detects sequences resulting from internal priming or from alternative splicing, identical sequences or sequences with several frameshifts. This automatic analysis serves as a basis for manual selection of the sequences.

#### b) Manual sequence selection

Manual selection is carried out using automatically generated reports for each sequenced full length extended cDNA clone. During this manual procedures, a selection is operated between clones belonging to the same class as follows. ORF sequences encoded by clones belonging to the same class are aligned and compared. If the homology between nucleotidic sequences of clones belonging to the same class is more than 90% over 30 nucleotide stretches or if the homology between amino acid sequences of clones belonging to the same class is more than 80% over 20 amino acid stretches, then the clones are considered as being identical. The chosen ORF is the best one according to the criteria mentioned below. If the nucleotide and amino acid homologies are less than 90% and 80% respectively, the clones are said to encode distinct proteins which can be both selected if they contain sequences of interest.

Selection of full length extended cDNA clones encoding sequences of interest is performed using the following criteria. Structural parameters (initial tag, polyadenylation

site and signal) are first checked. Then, homologies with known nucleic acids and proteins are examined in order to determine whether the clone sequence match a known nucleic/proteic sequence and, in the latter case, its covering rate and the date at which the sequence became public. If there is no extensive match with sequences other than ESTs or genomic DNA, or if the clone sequence brings substantial new information, such as encoding a protein resulting from alternative slicing of an mRNA coding for an already known protein, the sequence is kept. Examples of such cloned full length extended cDNAs containing sequences of interest are described in Example 28. Sequences resulting from chimera or double inserts as assessed by homology to other sequences are discarded during this procedure.

### EXAMPLE 28

#### Cloning and Sequencing of Extended cDNAs

The procedure described in Example 27 above was used to obtain the extended cDNAs of the present invention. Using this approach, the full length cDNA of SEQ ID NO:17 was obtained. This cDNA falls into the "EST-ext" category described above and encodes the signal peptide MKKVLLLITAILAVAVG (SEQ ID NO: 18) having a von Heijne score of 8.2.

The full length cDNA of SEQ ID NO: 19 was also obtained using this procedure. This cDNA falls into the "EST-ext" category described above and encodes the signal peptide MWWFQQGLSFLPSALVIWTS (SEQ ID NO:20) having a von Heijne score of 5.5.

Another full length cDNA obtained using the procedure described above has the sequence of SEQ ID NO:21. This cDNA, falls into the "EST-ext" category described above and encodes the signal peptide MVLTTLP SANSANSPVNMPTTGPNLSYASSA LSPCLT (SEQ ID NO:22) having a von Heijne score of 5.9.

The above procedure was also used to obtain a full length cDNA having the sequence of SEQ ID NO:23. This cDNA falls into the "EST-ext" category described above and encodes the signal peptide ILSTVTALTFAXA (SEQ ID NO:24) having a von Heijne score of 5.5.

The full length cDNA of SEQ ID NO:25 was also obtained using this procedure. This cDNA falls into the "new" category described above and encodes a signal peptide LVLTLCTLPLAVA (SEQ ID NO:26) having a von Heijne score of 10.1.

5 The full length cDNA of SEQ ID NO:27 was also obtained using this procedure. This cDNA falls into the "new" category described above and encodes a signal peptide LWLLFFLVTAIHA (SEQ ID NO:28) having a von Heijne score of 10.7.

10 The above procedures were also used to obtain the extended cDNAs of the present invention. 5' ESTs expressed in a variety of tissues were obtained as described above. The appended sequence listing provides the tissues from which the extended cDNAs were obtained. It will be appreciated that the extended cDNAs may also be expressed in tissues other than the tissue listed in the sequence listing.

15 5' ESTs obtained as described above were used to obtain extended cDNAs having the sequences of SEQ ID NOs: 40-140 and 242-377. Table IV provides the sequence identification numbers of the extended cDNAs of the present invention, the locations of the full coding sequences in SEQ ID NOs: 40-140 and 242-377 (i.e. the nucleotides encoding both the signal peptide and the mature protein, listed under the heading FCS location in Table IV), the locations of the nucleotides in SEQ ID NOs: 40-140 and 242-377 which encode the signal peptides (listed under the heading SigPep Location in Table IV), the locations of the nucleotides in SEQ ID NOs: 40-140 and 242-377 which encode the mature proteins generated by cleavage of the signal peptides (listed under the heading Mature Polypeptide Location in Table IV), the locations in SEQ ID NOs: 40-140 and 242-377 of stop codons (listed under the heading Stop Codon Location in Table IV), the locations in SEQ ID NOs: 40-140 and 242-377 of polyA signals (listed under the heading Poly A Signal Location in Table IV) and the locations of polyA sites (listed under the heading Poly A Site Location in Table IV).

20

25

The polypeptides encoded by the extended cDNAs were screened for the presence of known structural or functional motifs or for the presence of signatures, small amino acid sequences which are well conserved amongst the members of a protein family. The conserved regions have been used to derive consensus patterns or matrices included in the

PROSITE data bank, in particular in the file prosite.dat (Release 13.0 of November 1995, located at <http://expasy.hcuge.ch/sprot/prosite.html>. Prosite\_convert and prosite\_scan programs ([http://ulrec3.unil.ch/ftpserveur/prosite\\_scan](http://ulrec3.unil.ch/ftpserveur/prosite_scan)) were used to find signatures on the extended cDNAs.

5 For each pattern obtained with the prosite\_convert program from the prosite.dat file, the accuracy of the detection on a new protein sequence has been tested by evaluating the frequency of irrelevant hits on the population of human secreted proteins included in the data bank SWISSPROT. The ratio between the number of hits on shuffled proteins (with a window size of 20 amino acids) and the number of hits on native (unshuffled)  
10 proteins was used as an index. Every pattern for which the ration was greater than 20% (one hit on shuffled proteins for 5 hits on native proteins) was skipped during the search with prosite\_scan. The program used to shuffle protein sequences (db\_shuffled) and the program used to determine the statistics for each pattern in the protein data banks (prosite\_statistics) are available on the ftp site [http://ulrec3.unil.ch/ftpserveur/prosite\\_scan](http://ulrec3.unil.ch/ftpserveur/prosite_scan).

15 Table V lists the sequence identification numbers of the polypeptides of SEQ ID NOs: 141-241 and 378-513, the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the full length polypeptide (second column), the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the signal peptides (third column), and the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-  
20 513 in the mature polypeptide created by cleaving the signal peptide from the full length polypeptide (fourth column).

The nucleotide sequences of the sequences of SEQ ID NOs: 40-140 and 242-377 and the amino acid sequences encoded by SEQ ID NOs: 40-140 and 242-377 (i.e. amino acid sequences of SEQ ID NOs: 141-241 and 378-513) are provided in the appended  
25 sequence listing. In some instances, the sequences are preliminary and may include some incorrect or ambiguous sequences or amino acids. The sequences of SEQ ID NOs: 40-140 and 242-377 can readily be screened for any errors therein and any sequence ambiguities can be resolved by resequencing a fragment containing such errors or ambiguities on both strands. Nucleic acid fragments for resolving sequencing errors or ambiguities may be



obtained from the deposited clones or can be isolated using the techniques described herein. Resolution of any such ambiguities or errors may be facilitated by using primers which hybridize to sequences located close to the ambiguous or erroneous sequences. For example, the primers may hybridize to sequences within 50-75 bases of the ambiguity or error. Upon resolution of an error or ambiguity, the corresponding corrections can be made in the protein sequences encoded by the DNA containing the error or ambiguity. For example, in the sequences of the present invention, ambiguities in the sequence of SEQ ID NO: 131 were resolved. The amino acid sequence of the protein encoded by a particular clone can also be determined by expression of the clone in a suitable host cell, collecting the protein, and determining its sequence.

For each amino acid sequence, Applicants have identified what they have determined to be the reading frame best identifiable with sequence information available at the time of filing. Some of the amino acid sequences may contain "Xaa" designators. These "Xaa" designators indicate either (1) a residue which cannot be identified because of nucleotide sequence ambiguity or (2) a stop codon in the determined sequence where Applicants believe one should not exist (if the sequence were determined more accurately).

Cells containing the extended cDNAs (SEQ ID NOs: 40-140 and 242-377) of the present invention in the vector pED6dpc2, are maintained in permanent deposit by the inventors at Genset, S.A., 24 Rue Royale, 75008 Paris, France.

Pools of cells containing the extended cDNAs (SEQ ID NOs: 40-140 and 242-377), from which cells containing a particular polynucleotide are obtainable, were deposited with the American Type Culture Collection, 10801 University Blvd., Manassas, VA 20110-2209 or the European Collection of Cell Cultures, Vaccine Research and Production Laboratory, Public Health Laboratory Service, Centre for Applied Microbiology and Research, Porton Down, Salisbury, Wiltshire SP4 0JG, United Kingdom. Each extended cDNA clone has been transfected into separate bacterial cells (E-coli) for this composite deposit. Table VI lists the deposit numbers of the clones of SEQ ID Nos: 40-140. One or more pools of cells containing the extended cDNAs of SEQ

ID Nos: 242-377, from which the cells containing a particular polynucleotide is obtainable, will be deposited with the European Collection of Cell Cultures, Vaccine Research and Production Laboratory, Public Health Laboratory Service, Centre for Applied Microbiology and Research, Porton Down, Salisbury, Wiltshire SP4 OJG, United Kingdom and will be assigned ECACC deposit number XXXXXXXX. Table VII provides the internal designation number assigned to each SEQ ID NO. and indicates whether the sequence is a nucleic acid sequence or a protein sequence.

Each extended cDNA can be removed from the pED6dpc2 vector in which it was deposited by performing a NotI, PstI double digestion to produce the appropriate fragment for each clone. The proteins encoded by the extended cDNAs may also be expressed from the promoter in pED6dpc2.

Bacterial cells containing a particular clone can be obtained from the composite deposit as follows:

An oligonucleotide probe or probes should be designed to the sequence that is known for that particular clone. This sequence can be derived from the sequences provided herein, or from a combination of those sequences. The design of the oligonucleotide probe should preferably follow these parameters:

(a) It should be designed to an area of the sequence which has the fewest ambiguous bases ("N's"), if any;

(b) Preferably, the probe is designed to have a  $T_m$  of approx. 80°C (assuming 2 degrees for each A or T and 4 degrees for each G or C). However, probes having melting temperatures between 40 °C and 80 °C may also be used provided that specificity is not lost.

The oligonucleotide should preferably be labeled with  $\gamma$ -[ $^{32}\text{P}$ ]ATP (specific activity 6000 Ci/mmol) and T4 polynucleotide kinase using commonly employed techniques for labeling oligonucleotides. Other labeling techniques can also be used. Unincorporated label should preferably be removed by gel filtration chromatography or other established methods. The amount of radioactivity incorporated into the probe

should be quantified by measurement in a scintillation counter. Preferably, specific activity of the resulting probe should be approximately  $4 \times 10^6$  dpm/pmole.

5 The bacterial culture containing the pool of full-length clones should preferably be thawed and 100  $\mu$ l of the stock used to inoculate a sterile culture flask containing 25 ml of sterile L-broth containing ampicillin at 100  $\mu$ g/ml. The culture should preferably be grown to saturation at 37°C, and the saturated culture should preferably be diluted in fresh L-broth. Aliquots of these dilutions should preferably be plated to determine the dilution and volume which will yield approximately 5000 distinct and well-separated colonies on solid bacteriological media containing L-broth containing ampicillin at 100  $\mu$ g/ml and 10 agar at 1.5% in a 150 mm petri dish when grown overnight at 37°C. Other known methods of obtaining distinct, well-separated colonies can also be employed.

Standard colony hybridization procedures should then be used to transfer the colonies to nitrocellulose filters and lyse, denature and bake them.

15 The filter is then preferably incubated at 65°C for 1 hour with gentle agitation in 6X SSC (20X stock is 175.3 g NaCl/liter, 88.2 g Na citrate/liter, adjusted to pH 7.0 with NaOH) containing 0.5% SDS, 100  $\mu$ g/ml of yeast RNA, and 10 mM EDTA (approximately 10 mL per 150 mm filter). Preferably, the probe is then added to the hybridization mix at a concentration greater than or equal to  $1 \times 10^6$  dpm/mL. The filter is then preferably incubated at 65°C with gentle agitation overnight. The filter is then 20 preferably washed in 500 mL of 2X SSC/0.1% SDS at room temperature with gentle shaking for 15 minutes. A third wash with 0.1X SSC/0.5% SDS at 65°C for 30 minutes to 1 hour is optional. The filter is then preferably dried and subjected to autoradiography for sufficient time to visualize the positives on the X-ray film. Other known hybridization methods can also be employed.

25 The positive colonies are picked, grown in culture, and plasmid DNA isolated using standard procedures. The clones can then be verified by restriction analysis, hybridization analysis, or DNA sequencing.

The plasmid DNA obtained using these procedures may then be manipulated using standard cloning techniques familiar to those skilled in the art. Alternatively, a PCR can

be done with primers designed at both ends of the extended cDNA insertion. For example, a PCR reaction may be conducted using a primer having the sequence GGCCATACTTGAGTGAC (SEQ ID NO:38) and a primer having the sequence ATATAGACAAACGCACACC (SEQ. ID. NO:39). The PCR product which corresponds to the extended cDNA can then be manipulated using standard cloning techniques familiar to those skilled in the art.

In addition to PCR based methods for obtaining extended cDNAs, traditional hybridization based methods may also be employed. These methods may also be used to obtain the genomic DNAs which encode the mRNAs from which the 5' ESTs were derived, mRNAs corresponding to the extended cDNAs, or nucleic acids which are homologous to extended cDNAs or 5' ESTs. Example 29 below provides an example of such methods.

#### EXAMPLE 29

##### Methods for Obtaining Extended cDNAs or Nucleic Acids Homologous to Extended cDNAs or 5' ESTs

A full length cDNA library can be made using the strategies described in Examples 13, 14, 15, and 16 above by replacing the random nonamer used in Example 14 with an oligo-dT primer. For instance, the oligonucleotide of SEQ ID NO:14 may be used.

Alternatively, a cDNA library or genomic DNA library may be obtained from a commercial source or made using techniques familiar to those skilled in the art. The library includes cDNAs which are derived from the mRNA corresponding to a 5' EST or which have homology to an extended cDNA or 5' EST. The cDNA library or genomic DNA library is hybridized to a detectable probe comprising at least 10 consecutive nucleotides from the 5' EST or extended cDNA using conventional techniques. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST or extended cDNA. More preferably, the probe comprises at least 20-30 consecutive nucleotides from the 5' EST or extended cDNA. In some embodiments, the probe comprises at least 30 nucleotides from the 5' EST or extended cDNA. In other

embodiments, the probe comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the 5' EST or extended cDNA.

Techniques for identifying cDNA clones in a cDNA library which hybridize to a given probe sequence are disclosed in Sambrook et al., Molecular Cloning: A Laboratory Manual 2d Ed., Cold Spring Harbor Laboratory Press, 1989. The same techniques may be used to isolate genomic DNAs.

Briefly, cDNA or genomic DNA clones which hybridize to the detectable probe are identified and isolated for further manipulation as follows. A probe comprising at least 10 consecutive nucleotides from the 5' EST or extended cDNA is labeled with a detectable label such as a radioisotope or a fluorescent molecule. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST or extended cDNA. More preferably, the probe comprises 20-30 consecutive nucleotides from the 5' EST or extended cDNA. In some embodiments, the probe comprises more than 30 nucleotides from the 5' EST or extended cDNA. In some embodiments, the probe comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the 5' EST or extended cDNA.

Techniques for labeling the probe are well known and include phosphorylation with polynucleotide kinase, nick translation, in vitro transcription, and non-radioactive techniques. The cDNAs or genomic DNAs in the library are transferred to a nitrocellulose or nylon filter and denatured. After incubation of the filter with a blocking solution, the filter is contacted with the labeled probe and incubated for a sufficient amount of time for the probe to hybridize to cDNAs or genomic DNAs containing a sequence capable of hybridizing to the probe.

By varying the stringency of the hybridization conditions used to identify extended cDNAs or genomic DNAs which hybridize to the detectable probe, extended cDNAs having different levels of homology to the probe can be identified and isolated. To identify extended cDNAs or genomic DNAs having a high degree of homology to the probe sequence, the melting temperature of the probe may be calculated using the following formulas:

For probes between 14 and 70 nucleotides in length the melting temperature ( $T_m$ ) is calculated using the formula:  $T_m = 81.5 + 16.6(\log [Na^+]) + 0.41(\text{fraction G+C}) - (600/N)$  where N is the length of the probe.

5 If the hybridization is carried out in a solution containing formamide, the melting temperature may be calculated using the equation  $T_m = 81.5 + 16.6(\log [Na^+]) + 0.41(\text{fraction G+C}) - (0.63\% \text{ formamide}) - (600/N)$  where N is the length of the probe.

Prehybridization may be carried out in 6X SSC, 5X Denhardt's reagent, 0.5% SDS, 100 $\mu$ g denatured fragmented salmon sperm DNA or 6X SSC, 5X Denhardt's reagent, 0.5% SDS, 100 $\mu$ g denatured fragmented salmon sperm DNA, 50% formamide. 10 The formulas for SSC and Denhardt's solutions are listed in Sambrook et al., supra.

Hybridization is conducted by adding the detectable probe to the prehybridization solutions listed above. Where the probe comprises double stranded DNA, it is denatured before addition to the hybridization solution. The filter is contacted with the hybridization solution for a sufficient period of time to allow the probe to hybridize to extended cDNAs or genomic DNAs containing sequences complementary thereto or homologous thereto. 15 For probes over 200 nucleotides in length, the hybridization may be carried out at 15-25°C below the  $T_m$ . For shorter probes, such as oligonucleotide probes, the hybridization may be conducted at 15-25°C below the  $T_m$ . Preferably, for hybridizations in 6X SSC, the hybridization is conducted at approximately 68°C. Preferably, for hybridizations in 50% 20 formamide containing solutions, the hybridization is conducted at approximately 42°C.

All of the foregoing hybridizations would be considered to be under "stringent" conditions. Following hybridization, the filter is washed in 2X SSC, 0.1% SDS at room temperature for 15 minutes. The filter is then washed with 0.1X SSC, 0.5% SDS at room 25 temperature for 30 minutes to 1 hour. Thereafter, the solution is washed at the hybridization temperature in 0.1X SSC, 0.5% SDS. A final wash is conducted in 0.1X SSC at room temperature.

Extended cDNAs, nucleic acids homologous to extended cDNAs or 5' ESTs, or genomic DNAs which have hybridized to the probe are identified by autoradiography or other conventional techniques.

5 The above procedure may be modified to identify extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs having decreasing levels of homology to the probe sequence. For example, to obtain extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs of decreasing homology to the detectable probe, less stringent conditions may be used. For example, the hybridization temperature may be decreased in increments of 5°C from 68°C to 42°C in a hybridization  
10 buffer having a Na<sup>+</sup> concentration of approximately 1M. Following hybridization, the filter may be washed with 2X SSC, 0.5% SDS at the temperature of hybridization. These conditions are considered to be "moderate" conditions above 50°C and "low" conditions below 50°C.

15 Alternatively, the hybridization may be carried out in buffers, such as 6X SSC, containing formamide at a temperature of 42°C. In this case, the concentration of formamide in the hybridization buffer may be reduced in 5% increments from 50% to 0% to identify clones having decreasing levels of homology to the probe. Following hybridization, the filter may be washed with 6X SSC, 0.5% SDS at 50°C. These conditions are considered to be "moderate" conditions above 25% formamide and "low"  
20 conditions below 25% formamide.

Extended cDNAs, nucleic acids homologous to extended cDNAs, or genomic DNAs which have hybridized to the probe are identified by autoradiography.

25 If it is desired to obtain nucleic acids homologous to extended cDNAs, such as allelic variants thereof or nucleic acids encoding proteins related to the proteins encoded by the extended cDNAs, the level of homology between the hybridized nucleic acid and the extended cDNA or 5' EST used as the probe may readily be determined. To determine the level of homology between the hybridized nucleic acid and the extended cDNA or 5'EST from which the probe was derived, the nucleotide sequences of the hybridized nucleic acid and the extended cDNA or 5'EST from which the probe was derived are

compared. For example, using the above methods, nucleic acids having at least 95% nucleic acid homology to the extended cDNA or 5'EST from which the probe was derived may be obtained and identified. Similarly, by using progressively less stringent hybridization conditions one can obtain and identify nucleic acids having at least 90%, at least 85%, at least 80% or at least 75% homology to the extended cDNA or 5'EST from which the probe was derived. The level of homology between the hybridized nucleic acid and the extended cDNA or 5' EST used as the probe may be further determined using BLAST2N; parameters may be adapted depending on the sequence length and degree of homology studied. In such comparisons, the default parameters or the parameters listed in Tables II and III may be used.

To determine whether a clone encodes a protein having a given amount of homology to the protein encoded by the extended cDNA or 5' EST, the amino acid sequence encoded by the extended cDNA or 5' EST is compared to the amino acid sequence encoded by the hybridizing nucleic acid. Homology is determined to exist when an amino acid sequence in the extended cDNA or 5' EST is closely related to an amino acid sequence in the hybridizing nucleic acid. A sequence is closely related when it is identical to that of the extended cDNA or 5' EST or when it contains one or more amino acid substitutions therein in which amino acids having similar characteristics have been substituted for one another. Using the above methods, one can obtain nucleic acids encoding proteins having at least 95%, at least 90%, at least 85%, at least 80% or at least 75% homology to the proteins encoded by the extended cDNA or 5'EST from which the probe was derived. Using the above methods and algorithms such as FASTA with parameters depending on the sequence length and degree of homology studied the level of homology may be determined. In determining the level of homology using FASTA, the default parameters or the parameters listed in Tables II or III may be used.

Alternatively, extended cDNAs may be prepared by obtaining mRNA from the tissue, cell, or organism of interest using mRNA preparation procedures utilizing poly A selection procedures or other techniques known to those skilled in the art. A first primer



capable of hybridizing to the poly A tail of the mRNA is hybridized to the mRNA and a reverse transcription reaction is performed to generate a first cDNA strand.

5 The first cDNA strand is hybridized to a second primer containing at least 10 consecutive nucleotides of the sequences of the 5' EST for which an extended cDNA is desired. Preferably, the primer comprises at least 12, 15, or 17 consecutive nucleotides from the sequences of the 5' EST. More preferably, the primer comprises 20-30 consecutive nucleotides from the sequences of the 5' EST. In some embodiments, the primer comprises more than 30 nucleotides from the sequences of the 5' EST. If it is desired to obtain extended cDNAs containing the full protein coding sequence, including  
10 the authentic translation initiation site, the second primer used contains sequences located upstream of the translation initiation site. The second primer is extended to generate a second cDNA strand complementary to the first cDNA strand. Alternatively, RTPCR may be performed as described above using primers from both ends of the cDNA to be obtained.

15 Extended cDNAs containing 5' fragments of the mRNA may be prepared by contacting an mRNA comprising the sequence of the 5' EST for which an extended cDNA is desired with a primer comprising at least 10 consecutive nucleotides of the sequences complementary to the 5' EST, hybridizing the primer to the mRNAs, and reverse transcribing the hybridized primer to make a first cDNA strand from the mRNAs.  
20 Preferably, the primer comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST. More preferably, the primer comprises 20-30 consecutive nucleotides from the 5' EST.

Thereafter, a second cDNA strand complementary to the first cDNA strand is synthesized. The second cDNA strand may be made by hybridizing a primer  
25 complementary to sequences in the first cDNA strand to the first cDNA strand and extending the primer to generate the second cDNA strand.

The double stranded extended cDNAs made using the methods described above are isolated and cloned. The extended cDNAs may be cloned into vectors such as

plasmids or viral vectors capable of replicating in an appropriate host cell. For example, the host cell may be a bacterial, mammalian, avian, or insect cell.

Techniques for isolating mRNA, reverse transcribing a primer hybridized to mRNA to generate a first cDNA strand, extending a primer to make a second cDNA strand complementary to the first cDNA strand, isolating the double stranded cDNA and cloning the double stranded cDNA are well known to those skilled in the art and are described in Current Protocols in Molecular Biology, John Wiley 503 Sons, Inc. 1997 and Sambrook et al. Molecular Cloning: A Laboratory Manual, Second Edition, Cold Spring Harbor Laboratory Press, 1989.

Alternatively, kits for obtaining full length cDNAs, such as the GeneTrapper (Cat. No. 10356-020, Gibco, BRL), may be used for obtaining full length cDNAs or extended cDNAs. In this approach, full length or extended cDNAs are prepared from mRNA and cloned into double stranded phagemids. The cDNA library in the double stranded phagemids is then rendered single stranded by treatment with an endonuclease, such as the Gene II product of the phage F1, and Exonuclease III as described in the manual accompanying the GeneTrapper kit. A biotinylated oligonucleotide comprising the sequence of a 5' EST, or a fragment containing at least 10 nucleotides thereof, is hybridized to the single stranded phagemids. Preferably, the fragment comprises at least 12, 15, or 17 consecutive nucleotides from the 5' EST. More preferably, the fragment comprises 20-30 consecutive nucleotides from the 5' EST. In some procedures, the fragment may comprise more than 30 consecutive nucleotides from the 5' EST. For example, the fragment may comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the 5' EST.

Hybrids between the biotinylated oligonucleotide and phagemids having inserts containing the 5' EST sequence are isolated by incubating the hybrids with streptavidin coated paramagnetic beads and retrieving the beads with a magnet. Thereafter, the resulting phagemids containing the 5' EST sequence are released from the beads and converted into double stranded DNA using a primer specific for the 5' EST sequence. The

resulting double stranded DNA is transformed into bacteria. Extended cDNAs containing the 5' EST sequence are identified by colony PCR or colony hybridization.

5 A plurality of extended cDNAs containing full length protein coding sequences or sequences encoding only the mature protein remaining after the signal peptide is cleaved may be provided as cDNA libraries for subsequent evaluation of the encoded proteins or use in diagnostic assays as described below.

#### **IV. Expression of Proteins Encoded by Extended cDNAs Isolated Using 5' ESTs**

10 Extended cDNAs containing the full protein coding sequences of their corresponding mRNAs or portions thereof, such as cDNAs encoding the mature protein, may be used to express the secreted proteins or portions thereof which they encode as described in Example 30 below. If desired, the extended cDNAs may contain the sequences encoding the signal peptide to facilitate secretion of the expressed protein. It will be appreciated that a plurality of extended cDNAs containing the full protein coding sequences or portions thereof may be simultaneously cloned into expression vectors to create an expression library for analysis of the encoded proteins as described below.

#### **EXAMPLE 30**

##### Expression of the Proteins Encoded by Extended cDNAs or Portions Thereof

20 To express the proteins encoded by the extended cDNAs or portions thereof, nucleic acids containing the coding sequence for the proteins or portions thereof to be expressed are obtained as described in Examples 27-29 and cloned into a suitable expression vector. If desired, the nucleic acids may contain the sequences encoding the signal peptide to facilitate secretion of the expressed protein. For example, the nucleic acid may comprise the sequence of one of SEQ ID NOs: 40-140 and 242-377 listed in Table IV and in the accompanying sequence listing. Alternatively, the nucleic acid may  
25 comprise those nucleotides which make up the full coding sequence of one of the sequences of SEQ ID NOs: 40-140 and 242-377 as defined in Table IV above.

It will be appreciated that should the extent of the full coding sequence (i.e. the sequence encoding the signal peptide and the mature protein resulting from cleavage of

the signal peptide) differ from that listed in Table IV as a result of a sequencing error, reverse transcription or amplification error, mRNA splicing, post-translational modification of the encoded protein, enzymatic cleavage of the encoded protein, or other biological factors, one skilled in the art would be readily able to identify the extent of the full coding sequences in the sequences of SEQ ID NOs. 40-140 and 242-377. For example, the sequence of SEQ ID NO: 115 represents an alternatively spliced transcript of a previously identified mRNA. Accordingly, the scope of any claims herein relating to nucleic acids containing the full coding sequence of one of SEQ ID NOs. 40-140 and 242-377 is not to be construed as excluding any readily identifiable variations from or equivalents to the full coding sequences listed in Table IV. Similarly, should the extent of the full length polypeptides differ from those indicated in Table V as a result of any of the preceding factors, the scope of claims relating to polypeptides comprising the amino acid sequence of the full length polypeptides is not to be construed as excluding any readily identifiable variations from or equivalents to the sequences listed in Table V.

Alternatively, the nucleic acid used to express the protein or portion thereof may comprise those nucleotides which encode the mature protein (i.e. the protein created by cleaving the signal peptide off) encoded by one of the sequences of SEQ ID NOs: 40-140 and 242-377 as defined in Table IV above.

It will be appreciated that should the extent of the sequence encoding the mature protein differ from that listed in Table IV as a result of a sequencing error, reverse transcription or amplification error, mRNA splicing, post-translational modification of the encoded protein, enzymatic cleavage of the encoded protein, or other biological factors, one skilled in the art would be readily able to identify the extent of the sequence encoding the mature protein in the sequences of SEQ ID NOs. 40-140 and 242-377. Accordingly, the scope of any claims herein relating to nucleic acids containing the sequence encoding the mature protein encoded by one of SEQ ID Nos. 40-140 and 242-377 is not to be construed as excluding any readily identifiable variations from or equivalents to the sequences listed in Table IV. Thus, claims relating to nucleic acids containing the sequence encoding the mature protein encompass equivalents to the sequences listed in

Table IV, such as sequences encoding biologically active proteins resulting from post-translational modification, enzymatic cleavage, or other readily identifiable variations from or equivalents to the secreted proteins in addition to cleavage of the signal peptide. Similarly, should the extent of the mature polypeptides differ from those indicated in Table V as a result of any of the preceding factors, the scope of claims relating to polypeptides comprising the sequence of a mature protein included in the sequence of one of SEQ ID NOs. 141-241 and 378-513 is not to be construed as excluding any readily identifiable variations from or equivalents to the sequences listed in Table V. Thus, claims relating to polypeptides comprising the sequence of the mature protein encompass equivalents to the sequences listed in Table IV, such as biologically active proteins resulting from post-translational modification, enzymatic cleavage, or other readily identifiable variations from or equivalents to the secreted proteins in addition to cleavage of the signal peptide. It will also be appreciated that should the biologically active form of the polypeptides included in the sequence of one of SEQ ID NOs. 141-241 and 378-513 or the nucleic acids encoding the biologically active form of the polypeptides differ from those identified as the mature polypeptide in Table V or the nucleotides encoding the mature polypeptide in Table IV as a result of a sequencing error, reverse transcription or amplification error, mRNA splicing, post-translational modification of the encoded protein, enzymatic cleavage of the encoded protein, or other biological factors, one skilled in the art would be readily able to identify the amino acids in the biologically active form of the polypeptides and the nucleic acids encoding the biologically active form of the polypeptides. In such instances, the claims relating to polypeptides comprising the mature protein included in one of SEQ ID NOs. 141-241 and 378-513 or nucleic acids comprising the nucleotides of one of SEQ ID NOs. 40-140 and 242-377 encoding the mature protein shall not be construed to exclude any readily identifiable variations from the sequences listed in Table IV and Table V.

In some embodiments, the nucleic acid used to express the protein or portion thereof may comprise those nucleotides which encode the signal peptide encoded by one of the sequences of SEQ ID NOs: 40-140 and 242-377 as defined in Table IV above.

It will be appreciated that should the extent of the sequence encoding the signal peptide differ from that listed in Table IV as a result of a sequencing error, reverse transcription or amplification error, mRNA splicing, post-translational modification of the encoded protein, enzymatic cleavage of the encoded protein, or other biological factors, one skilled in the art would be readily able to identify the extent of the sequence encoding the signal peptide in the sequences of SEQ ID NOs. 40-140 and 242-377. Accordingly, the scope of any claims herein relating to nucleic acids containing the sequence encoding the signal peptide encoded by one of SEQ ID Nos. 40-140 and 242-377 is not to be construed as excluding any readily identifiable variations from the sequences listed in Table IV. Similarly, should the extent of the signal peptides differ from those indicated in Table V as a result of any of the preceding factors, the scope of claims relating to polypeptides comprising the sequence of a signal peptide included in the sequence of one of SEQ ID NOs. 141-241 and 378-513 is not to be construed as excluding any readily identifiable variations from the sequences listed in Table V.

Alternatively, the nucleic acid may encode a polypeptide comprising at least 10 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In some embodiments, the nucleic acid may encode a polypeptide comprising at least 15 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In other embodiments, the nucleic acid may encode a polypeptide comprising at least 25 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513. In other embodiments, the nucleic acid may encode a polypeptide comprising at least 60, at least 75, at least 100 or more than 100 consecutive amino acids of one of the sequences of SEQ ID Nos: 141-241 and 378-513.

The nucleic acids inserted into the expression vectors may also contain sequences upstream of the sequences encoding the signal peptide, such as sequences which regulate expression levels or sequences which confer tissue specific expression.

The nucleic acid encoding the protein or polypeptide to be expressed is operably linked to a promoter in an expression vector using conventional cloning technology. The expression vector may be any of the mammalian, yeast, insect or bacterial expression

systems known in the art. Commercially available vectors and expression systems are available from a variety of suppliers including Genetics Institute (Cambridge, MA), Stratagene (La Jolla, California), Promega (Madison, Wisconsin), and Invitrogen (San Diego, California). If desired, to enhance expression and facilitate proper protein folding, the codon context and codon pairing of the sequence may be optimized for the particular expression organism in which the expression vector is introduced, as explained by Hatfield, et al., U.S. Patent No. 5,082,767, incorporated herein by this reference.

The following is provided as one exemplary method to express the proteins encoded by the extended cDNAs corresponding to the 5' ESTs or the nucleic acids described above. First, the methionine initiation codon for the gene and the poly A signal of the gene are identified. If the nucleic acid encoding the polypeptide to be expressed lacks a methionine to serve as the initiation site, an initiating methionine can be introduced next to the first codon of the nucleic acid using conventional techniques. Similarly, if the extended cDNA lacks a poly A signal, this sequence can be added to the construct by, for example, splicing out the Poly A signal from pSG5 (Stratagene) using BglII and SalI restriction endonuclease enzymes and incorporating it into the mammalian expression vector pXT1 (Stratagene). pXT1 contains the LTRs and a portion of the *gag* gene from Moloney Murine Leukemia Virus. The position of the LTRs in the construct allow efficient stable transfection. The vector includes the Herpes Simplex Thymidine Kinase promoter and the selectable neomycin gene. The extended cDNA or portion thereof encoding the polypeptide to be expressed is obtained by PCR from the bacterial vector using oligonucleotide primers complementary to the extended cDNA or portion thereof and containing restriction endonuclease sequences for Pst I incorporated into the 5' primer and BglII at the 5' end of the corresponding cDNA 3' primer, taking care to ensure that the extended cDNA is positioned in frame with the poly A signal. The purified fragment obtained from the resulting PCR reaction is digested with PstI, blunt ended with an exonuclease, digested with Bgl II, purified and ligated to pXT1, now containing a poly A signal and digested with BglII.

The ligated product is transfected into mouse NIH 3T3 cells using Lipofectin (Life Technologies, Inc., Grand Island, New York) under conditions outlined in the product specification. Positive transfectants are selected after growing the transfected cells in 600ug/ml G418 (Sigma, St. Louis, Missouri). Preferably the expressed protein is released  
5 into the culture medium, thereby facilitating purification.

Alternatively, the extended cDNAs may be cloned into pED6dpc2 as described above. The resulting pED6dpc2 constructs may be transfected into a suitable host cell, such as COS 1 cells. Methotrexate resistant cells are selected and expanded. Preferably, the protein expressed from the extended cDNA is released into the culture medium  
10 thereby facilitating purification.

Proteins in the culture medium are separated by gel electrophoresis. If desired, the proteins may be ammonium sulfate precipitated or separated based on size or charge prior to electrophoresis.

As a control, the expression vector lacking a cDNA insert is introduced into host  
15 cells or organisms and the proteins in the medium are harvested. The secreted proteins present in the medium are detected using techniques such as Coomassie or silver staining or using antibodies against the protein encoded by the extended cDNA. Coomassie and silver staining techniques are familiar to those skilled in the art.

Antibodies capable of specifically recognizing the protein of interest may be  
20 generated using synthetic 15-mer peptides having a sequence encoded by the appropriate 5' EST, extended cDNA, or portion thereof. The synthetic peptides are injected into mice to generate antibody to the polypeptide encoded by the 5' EST, extended cDNA, or portion thereof.

Secreted proteins from the host cells or organisms containing an expression vector  
25 which contains the extended cDNA derived from a 5' EST or a portion thereof are compared to those from the control cells or organism. The presence of a band in the medium from the cells containing the expression vector which is absent in the medium from the control cells indicates that the extended cDNA encodes a secreted protein. Generally, the band corresponding to the protein encoded by the extended cDNA will



have a mobility near that expected based on the number of amino acids in the open reading frame of the extended cDNA. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

5           Alternatively, if the protein expressed from the above expression vectors does not contain sequences directing its secretion, the proteins expressed from host cells containing an expression vector containing an insert encoding a secreted protein or portion thereof can be compared to the proteins expressed in host cells containing the expression vector without an insert. The presence of a band in samples from cells containing the expression  
10       vector with an insert which is absent in samples from cells containing the expression vector without an insert indicates that the desired protein or portion thereof is being expressed. Generally, the band will have the mobility expected for the secreted protein or portion thereof. However, the band may have a mobility different than that expected as a result of modifications such as glycosylation, ubiquitination, or enzymatic cleavage.

15           The protein encoded by the extended cDNA may be purified using standard immunochromatography techniques. In such procedures, a solution containing the secreted protein, such as the culture medium or a cell extract, is applied to a column having antibodies against the secreted protein attached to the chromatography matrix. The secreted protein is allowed to bind the immunochromatography column. Thereafter, the  
20       column is washed to remove non-specifically bound proteins. The specifically bound secreted protein is then released from the column and recovered using standard techniques.

          If antibody production is not possible, the extended cDNA sequence or portion thereof may be incorporated into expression vectors designed for use in purification  
25       schemes employing chimeric polypeptides. In such strategies the coding sequence of the extended cDNA or portion thereof is inserted in frame with the gene encoding the other half of the chimera. The other half of the chimera may be  $\beta$ -globin or a nickel binding polypeptide encoding sequence. A chromatography matrix having antibody to  $\beta$ -globin or nickel attached thereto is then used to purify the chimeric protein. Protease cleavage sites

may be engineered between the  $\beta$ -globin gene or the nickel binding polypeptide and the extended cDNA or portion thereof. Thus, the two polypeptides of the chimera may be separated from one another by protease digestion.

One useful expression vector for generating  $\beta$ -globin chimerics is pSG5 (Stratagene), which encodes rabbit  $\beta$ -globin. Intron II of the rabbit  $\beta$ -globin gene facilitates splicing of the expressed transcript, and the polyadenylation signal incorporated into the construct increases the level of expression. These techniques as described are well known to those skilled in the art of molecular biology. Standard methods are published in methods texts such as Davis et al., (**Basic Methods in Molecular Biology**, L.G. Davis, M.D. Dibner, and J.F. Battey, ed., Elsevier Press, NY, 1986) and many of the methods are available from Stratagene, Life Technologies, Inc., or Promega. Polypeptide may additionally be produced from the construct using in vitro translation systems such as the In vitro Express™ Translation Kit (Stratagene).

Following expression and purification of the secreted proteins encoded by the 5' ESTs, extended cDNAs, or fragments thereof, the purified proteins may be tested for the ability to bind to the surface of various cell types as described in Example 31 below. It will be appreciated that a plurality of proteins expressed from these cDNAs may be included in a panel of proteins to be simultaneously evaluated for the activities specifically described below, as well as other biological roles for which assays for determining activity are available.

### EXAMPLE 31

#### Analysis of Secreted Proteins to Determine Whether they Bind to the Cell Surface

The proteins encoded by the 5' ESTs, extended cDNAs, or fragments thereof are cloned into expression vectors such as those described in Example 30. The proteins are purified by size, charge, immunochromatography or other techniques familiar to those skilled in the art. Following purification, the proteins are labeled using techniques known to those skilled in the art. The labeled proteins are incubated with cells or cell lines derived from a variety of organs or tissues to allow the proteins to bind to any receptor present on the cell surface. Following the incubation, the cells are washed to remove non-

specifically bound protein. The labeled proteins are detected by autoradiography. Alternatively, unlabeled proteins may be incubated with the cells and detected with antibodies having a detectable label, such as a fluorescent molecule, attached thereto.

5 Specificity of cell surface binding may be analyzed by conducting a competition analysis in which various amounts of unlabeled protein are incubated along with the labeled protein. The amount of labeled protein bound to the cell surface decreases as the amount of competitive unlabeled protein increases. As a control, various amounts of an unlabeled protein unrelated to the labeled protein is included in some binding reactions. The amount of labeled protein bound to the cell surface does not decrease in binding  
10 reactions containing increasing amounts of unrelated unlabeled protein, indicating that the protein encoded by the cDNA binds specifically to the cell surface.

As discussed above, secreted proteins have been shown to have a number of important physiological effects and, consequently, represent a valuable therapeutic resource. The secreted proteins encoded by the extended cDNAs or portions thereof made  
15 according to Examples 27-29 may be evaluated to determine their physiological activities as described below.

### EXAMPLE 32

#### Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Cytokine, Cell Proliferation or Cell Differentiation Activity

20 As discussed above, secreted proteins may act as cytokines or may affect cellular proliferation or differentiation. Many protein factors discovered to date, including all known cytokines, have exhibited activity in one or more factor dependent cell proliferation assays, and hence the assays serve as a convenient confirmation of cytokine activity. The activity of a protein of the present invention is evidenced by any one of a  
25 number of routine factor dependent cell proliferation assays for cell lines including, without limitation, 32D, DA2, DA1G, T10, B9, B9/11, BaF3, MC9/G, M+ (preB M+), 2E8, RB5, DA1, 123, T1165, HT2, CTLL2, TF-1, Mo7c and CMK. The proteins encoded by the above extended cDNAs or portions thereof may be evaluated for their ability to regulate T cell or thymocyte proliferation in assays such as those described

above or in the following references, which are incorporated herein by reference:  
**Current Protocols in Immunology**, Ed. by J.E. Coligan et al., Greene Publishing  
Associates and Wiley-Interscience; Takai et al. **J. Immunol.** 137:3494-3500, 1986.  
Bertagnolli et al. **J. Immunol.** 145:1706-1712, 1990. Bertagnolli et al., **Cellular**  
5 **Immunology** 133:327-341, 1991. Bertagnolli, et al. **J. Immunol.** 149:3778-3783, 1992;  
Bowman et al., **J. Immunol.** 152:1756-1761, 1994.

In addition, numerous assays for cytokine production and/or the proliferation of  
spleen cells, lymph node cells and thymocytes are known. These include the techniques  
disclosed in **Current Protocols in Immunology**. J.E. Coligan et al. Eds., Vol 1 pp.  
10 3.12.1-3.12.14 John Wiley and Sons, Toronto. 1994; and Schreiber, R.D. **Current**  
**Protocols in Immunology**, *supra* Vol 1 pp. 6.8.1-6.8.8, John Wiley and Sons, Toronto.  
1994.

The proteins encoded by the cDNAs may also be assayed for the ability to regulate  
the proliferation and differentiation of hematopoietic or lymphopoietic cells. Many assays  
15 for such activity are familiar to those skilled in the art, including the assays in the  
following references, which are incorporated herein by reference: Bottomly, K., Davis,  
L.S. and Lipsky, P.E., Measurement of Human and Murine Interleukin 2 and Interleukin  
4, **Current Protocols in Immunology**, J.E. Coligan et al. Eds. Vol 1 pp. 6.3.1-6.3.12,  
John Wiley and Sons, Toronto. 1991; deVries et al., **J. Exp. Med.** 173:1205-1211, 1991;  
20 Moreau et al., **Nature** 36:690-692, 1988; Greenberger et al., **Proc. Natl. Acad. Sci.**  
**U.S.A.** 80:2931-2938, 1983; Nordan, R., Measurement of Mouse and Human Interleukin  
6 **Current Protocols in Immunology**. J.E. Coligan et al. Eds. Vol 1 pp. 6.6.1-6.6.5, John  
Wiley and Sons, Toronto. 1991; Smith et al., **Proc. Natl. Acad. Sci. U.S.A.** 83:1857-  
1861, 1986; Bennett, F., Giannotti, J., Clark, S.C. and Turner, K.J., Measurement of  
25 Human Interleukin 11 **Current Protocols in Immunology**. J.E. Coligan et al. Eds. Vol 1  
pp. 6.15.1 John Wiley and Sons, Toronto. 1991; Ciarletta, A., Giannotti, J., Clark, S.C.  
and Turner, K.J., Measurement of Mouse and Human Interleukin 9 **Current Protocols in**  
**Immunology**. J.E. Coligan et al., Eds. Vol 1 pp. 6.13.1, John Wiley and Sons, Toronto.  
1991.

The proteins encoded by the cDNAs may also be assayed for their ability to regulate T-cell responses to antigens. Many assays for such activity are familiar to those skilled in the art, including the assays described in the following references, which are incorporated herein by reference: Chapter 3 (In Vitro Assays for Mouse Lymphocyte Function), Chapter 6 (Cytokines and Their Cellular Receptors) and Chapter 7, (Immunologic Studies in Humans) in **Current Protocols in Immunology**, J.E. Coligan et al. Eds. Greene Publishing Associates and Wiley-Interscience; Weinberger et al., **Proc. Natl. Acad. Sci. USA** 77:6091-6095, 1980; Weinberger et al., **Eur. J. Immun.** 11:405-411, 1981; Takai et al., **J. Immunol.** 137:3494-3500, 1986; Takai et al., **J. Immunol.** 140:508-512, 1988.

Those proteins which exhibit cytokine, cell proliferation, or cell differentiation activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which induction of cell proliferation or differentiation is beneficial. Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

### EXAMPLE 33

#### Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Activity as Immune System Regulators

The proteins encoded by the cDNAs may also be evaluated for their effects as immune regulators. For example, the proteins may be evaluated for their activity to influence thymocyte or splenocyte cytotoxicity. Numerous assays for such activity are familiar to those skilled in the art including the assays described in the following references, which are incorporated herein by reference: Chapter 3 (In Vitro Assays for Mouse Lymphocyte Function 3.1-3.19) and Chapter 7 (Immunologic studies in Humans) in **Current Protocols in Immunology**, J.E. Coligan et al. Eds, Greene Publishing Associates and Wiley-Interscience; Herrmann et al., **Proc. Natl. Acad. Sci. USA** 78:2488-2492, 1981; Herrmann et al., **J. Immunol.** 128:1968-1974, 1982; Handa et al., **J. Immunol.** 135:1564-1572, 1985; Takai et al., **J. Immunol.** 137:3494-3500, 1986; Takai

et al., **J. Immunol.** 140:508-512, 1988; Herrmann et al., **Proc. Natl. Acad. Sci. USA** 78:2488-2492, 1981; Herrmann et al., **J. Immunol.** 128:1968-1974, 1982; Handa et al., **J. Immunol.** 135:1564-1572, 1985; Takai et al., **J. Immunol.** 137:3494-3500, 1986; Bowman et al., **J. Virology** 61:1992-1998; Takai et al., **J. Immunol.** 140:508-512, 1988; Bertagnolli et al., **Cellular Immunology** 133:327-341, 1991; Brown et al., **J. Immunol.** 153:3079-3092, 1994.

The proteins encoded by the cDNAs may also be evaluated for their effects on T-cell dependent immunoglobulin responses and isotype switching. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Maliszewski, **J. Immunol.** 144:3028-3033, 1990; Mond, J.J. and Brunswick, M Assays for B Cell Function: *In vitro* Antibody Production, Vol 1 pp. 3.8.1-3.8.16 in **Current Protocols in Immunology**. J.E. Coligan et al Eds., John Wiley and Sons, Toronto. 1994.

The proteins encoded by the cDNAs may also be evaluated for their effect on immune effector cells, including their effect on Th1 cells and cytotoxic lymphocytes. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Chapter 3 (In Vitro Assays for Mouse Lymphocyte Function 3.1-3.19) and Chapter 7 (Immunologic Studies in Humans) in **Current Protocols in Immunology**, J.E. Coligan et al. Eds., Greene Publishing Associates and Wiley-Interscience; Takai et al., **J. Immunol.** 137:3494-3500, 1986; Takai et al.; **J. Immunol.** 140:508-512, 1988; Bertagnolli et al., **J. Immunol.** 149:3778-3783, 1992.

The proteins encoded by the cDNAs may also be evaluated for their effect on dendritic cell mediated activation of naive T-cells. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Guery et al., **J. Immunol.** 134:536-544, 1995; Inaba et al., **Journal of Experimental Medicine** 173:549-559, 1991; Macatonia et al., **Journal of Immunology** 154:5071-5079, 1995; Porgador et al., **Journal of Experimental Medicine** 182:255-260, 1995; Nair et al., **Journal of Virology** 67:4062-

4069, 1993; Huang et al., **Science** 264:961-965, 1994; Macatonia et al., **Journal of Experimental Medicine** 169:1255-1264, 1989; Bhardwaj et al., **Journal of Clinical Investigation** 94:797-807, 1994; and Inaba et al., **Journal of Experimental Medicine** 172:631-640, 1990.

5           The proteins encoded by the cDNAs may also be evaluated for their influence on the lifetime of lymphocytes. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Darzynkiewicz et al., **Cytometry** 13:795-808, 1992; Gorczyca et al., **Leukemia** 7:659-670, 1993; Gorczyca et al., **Cancer Research** 53:1945-10   1951, 1993; Itoh et al., **Cell** 66:233-243, 1991; Zacharchuk, **Journal of Immunology** 145:4037-4045, 1990; Zamai et al., **Cytometry** 14:891-897, 1993; Gorczyca et al., **International Journal of Oncology** 1:639-648, 1992.

          Assays for proteins that influence early steps of T-cell commitment and development include, without limitation, those described in: Antica et al., **Blood** 84:111-15   117, 1994; Fine et al., **Cellular immunology** 155:111-122, 1994; Galy et al., **Blood** 85:2770-2778, 1995; Toki et al., **Proc. Nat. Acad Sci. USA** 88:7548-7551, 1991.

          Those proteins which exhibit activity as immune system regulators activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of immune activity is beneficial. For example, the protein may be useful in the treatment of various immune deficiencies and disorders (including severe combined immunodeficiency (SCID)), e.g., in regulating (up or down) growth and proliferation of T and/or B lymphocytes, as well as effecting the cytolytic activity of NK cells and other cell populations. These immune deficiencies may be genetic or be caused by viral (e.g., HIV) as well as bacterial or fungal infections, or may result from autoimmune disorders. More 20   specifically, infectious diseases caused by viral, bacterial, fungal or other infection may be treatable using a protein of the present invention, including infections by HIV, hepatitis viruses, herpesviruses, mycobacteria, *Leishmania* spp., *malaria* spp. and various fungal infections such as candidiasis. Of course, in this regard, a protein of the present invention 25

may also be useful where a boost to the immune system generally may be desirable, i.e., in the treatment of cancer.

Autoimmune disorders which may be treated using a protein of the present invention include, for example, connective tissue disease, multiple sclerosis, systemic lupus erythematosus, rheumatoid arthritis, autoimmune pulmonary inflammation, Guillain-Barre syndrome, autoimmune thyroiditis, insulin dependent diabetes mellitis, myasthenia gravis, graft-versus-host disease and autoimmune inflammatory eye disease. Such a protein of the present invention may also to be useful in the treatment of allergic reactions and conditions, such as asthma (particularly allergic asthma) or other respiratory problems. Other conditions, in which immune suppression is desired (including, for example, organ transplantation), may also be treatable using a protein of the present invention.

Using the proteins of the invention it may also be possible to regulate immune responses, in a number of ways. Down regulation may be in the form of inhibiting or blocking an immune response already in progress or may involve preventing the induction of an immune response. The functions of activated T-cells may be inhibited by suppressing T cell responses or by inducing specific tolerance in T cells, or both. Immunosuppression of T cell responses is generally an active, non-antigen-specific, process which requires continuous exposure of the T cells to the suppressive agent. Tolerance, which involves inducing non-responsiveness or anergy in T cells, is distinguishable from immunosuppression in that it is generally antigen-specific and persists after exposure to the tolerizing agent has ceased. Operationally, tolerance can be demonstrated by the lack of a T cell response upon reexposure to specific antigen in the absence of the tolerizing agent.

Down regulating or preventing one or more antigen functions (including without limitation B lymphocyte antigen functions (such as, for example, B7)), e.g., preventing high level lymphokine synthesis by activated T cells, will be useful in situations of tissue, skin and organ transplantation and in graft-versus-host disease (GVHD). For example, blockage of T cell function should result in reduced tissue destruction in tissue



transplantation. Typically, in tissue transplants, rejection of the transplant is initiated through its recognition as foreign by T cells, followed by an immune reaction that destroys the transplant. The administration of a molecule which inhibits or blocks interaction of a B7 lymphocyte antigen with its natural ligand(s) on immune cells (such as a soluble, monomeric form of a peptide having B7-2 activity alone or in conjunction with a monomeric form of a peptide having an activity of another B lymphocyte antigen (e.g., B7-1, B7-3) or blocking antibody), prior to transplantation can lead to the binding of the molecule to the natural ligand(s) on the immune cells without transmitting the corresponding costimulatory signal. Blocking B lymphocyte antigen function in this matter prevents cytokine synthesis by immune cells, such as T cells, and thus acts as an immunosuppressant. Moreover, the lack of costimulation may also be sufficient to anergize the T cells, thereby inducing tolerance in a subject. Induction of long-term tolerance by B lymphocyte antigen-blocking reagents may avoid the necessity of repeated administration of these blocking reagents. To achieve sufficient immunosuppression or tolerance in a subject, it may also be necessary to block the function of a combination of B lymphocyte antigens.

The efficacy of particular blocking reagents in preventing organ transplant rejection or GVHD can be assessed using animal models that are predictive of efficacy in humans. Examples of appropriate systems which can be used include allogeneic cardiac grafts in rats and xenogeneic pancreatic islet cell grafts in mice, both of which have been used to examine the immunosuppressive effects of CTLA4Ig fusion proteins in vivo as described in Lenschow et al., Science 257:789-792 (1992) and Turka et al., Proc. Natl. Acad. Sci USA, 89:11102-11105 (1992). In addition, murine models of GVHD (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 846-847) can be used to determine the effect of blocking B lymphocyte antigen function in vivo on the development of that disease.

Blocking antigen function may also be therapeutically useful for treating autoimmune diseases. Many autoimmune disorders are the result of inappropriate activation of T cells that are reactive against self tissue and which promote the production

of cytokines and autoantibodies involved in the pathology of the diseases. Preventing the activation of autoreactive T cells may reduce or eliminate disease symptoms. Administration of reagents which block costimulation of T cells by disrupting receptor ligand interactions of B lymphocyte antigens can be used to inhibit T cell activation and prevent production of autoantibodies or T cell-derived cytokines which may be involved in the disease process. Additionally, blocking reagents may induce antigen-specific tolerance of autoreactive T cells which could lead to long-term relief from the disease. The efficacy of blocking reagents in preventing or alleviating autoimmune disorders can be determined using a number of well-characterized animal models of human autoimmune diseases. Examples include murine experimental autoimmune encephalitis, systemic lupus erythmatosis in MRL/pr/pr mice or NZB hybrid mice, murine autoimmune collagen arthritis, diabetes mellitus in OD mice and BB rats, and murine experimental myasthenia gravis (see Paul ed., Fundamental Immunology, Raven Press, New York, 1989, pp. 840-856).

Upregulation of an antigen function (preferably a B lymphocyte antigen function), as a means of up regulating immune responses, may also be useful in therapy. Upregulation of immune responses may be in the form of enhancing an existing immune response or eliciting an initial immune response. For example, enhancing an immune response through stimulating B lymphocyte antigen function may be useful in cases of viral infection. In addition, systemic viral diseases such as influenza, the common cold, and encephalitis might be alleviated by the administration of stimulatory form of B lymphocyte antigens systemically.

Alternatively, anti-viral immune responses may be enhanced in an infected patient by removing T cells from the patient, costimulating the T cells in vitro with viral antigen-pulsed APCs either expressing a peptide of the present invention or together with a stimulatory form of a soluble peptide of the present invention and reintroducing the in vitro activated T cells into the patient. The infected cells would now be capable of delivering a costimulatory signal to T cells in vivo, thereby activating the T cells.

In another application, up regulation or enhancement of antigen function (preferably B lymphocyte antigen function) may be useful in the induction of tumor immunity. Tumor cells (e.g., sarcoma, melanoma, lymphoma, leukemia, neuroblastoma, carcinoma) transfected with a nucleic acid encoding at least one peptide of the present invention can be administered to a subject to overcome tumor-specific tolerance in the subject. If desired, the tumor cell can be transfected to express a combination of peptides. For example, tumor cells obtained from a patient can be transfected ex vivo with an expression vector directing the expression of a peptide having B7-2-like activity alone, or in conjunction with a peptide having B7-1-like activity and/or B7-3-like activity. The transfected tumor cells are returned to the patient to result in expression of the peptides on the surface of the transfected cell. Alternatively, gene therapy techniques can be used to target a tumor cell for transfection in vivo.

The presence of the peptide of the present invention having the activity of a B lymphocyte antigen(s) on the surface of the tumor cell provides the necessary costimulation signal to T cells to induce a T cell mediated immune response against the transfected tumor cells. In addition, tumor cells which lack MHC class I or MHC class II molecules, or which fail to reexpress sufficient amounts of MHC class I or MHC class II molecules, can be transfected with nucleic acids encoding all or a portion of (e.g., a cytoplasmic-domain truncated portion) of an MHC class I  $\alpha$  chain protein and  $\beta_2$  macroglobulin protein or an MHC class II  $\alpha$  chain protein and an MHC class II  $\beta$  chain protein to thereby express MHC class I or MHC class II proteins on the cell surface. Expression of the appropriate class II or class II MHC in conjunction with a peptide having the activity of a B lymphocyte antigen (e.g., B7-1, B7-2, B7-3) induces a T cell mediated immune response against the transfected tumor cell. Optionally, a gene encoding an antisense construct which blocks expression of an MHC class II associated protein, such as the invariant chain, can also be cotransfected with a DNA encoding a peptide having the activity of a B lymphocyte antigen to promote presentation of tumor associated antigens and induce tumor specific immunity. Thus, the induction of a T cell mediated immune response in a human subject may be sufficient to overcome tumor-

specific tolerance in the subject. Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

5

#### EXAMPLE 34

##### Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Hematopoiesis Regulating Activity

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their hematopoiesis regulating activity. For example, the effect of the proteins on embryonic stem cell differentiation may be evaluated. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Johansson et al. **Cellular Biology** 15:141-151, 1995; Keller et al., **Molecular and Cellular Biology** 13:473-486, 1993; McClanahan et al., **Blood** 81:2903-2915, 1993.

15 The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their influence on the lifetime of stem cells and stem cell differentiation. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Freshney, M.G. Methylcellulose Colony Forming Assays, in **Culture of Hematopoietic Cells**. R.I. Freshney, et al. Eds. pp. 265-268, Wiley-Liss, Inc., New York, NY. 1994; Hirayama et al., **Proc. Natl. Acad. Sci. USA** 89:5907-5911, 1992; McNiece, I.K. and Briddell, R.A. Primitive Hematopoietic Colony Forming Cells with High Proliferative Potential, in **Culture of Hematopoietic Cells**. R.I. Freshney, et al. eds. Vol pp. 23-39, Wiley-Liss, Inc., New York, NY. 1994; Neben et al., **Experimental Hematology** 22:353-20 359, 1994; Ploemacher, R.E. Cobblestone Area Forming Cell Assay, In **Culture of Hematopoietic Cells**. R.I. Freshney, et al. Eds. pp. 1-21, Wiley-Liss, Inc., New York, NY. 1994; Spooncer, E., Dexter, M. and Allen, T. Long Term Bone Marrow Cultures in the Presence of Stromal Cells, in **Culture of Hematopoietic Cells**. R.I. Freshney, et al. Eds. pp. 163-179, Wiley-Liss, Inc., New York, NY. 1994; and Sutherland, H.J. Long

Term Culture Initiating Cell Assay, in **Culture of Hematopoietic Cells**. R.I. Freshney, et al. Eds. pp. 139-162, Wiley-Liss, Inc., New York, NY. 1994.

Those proteins which exhibit hematopoiesis regulatory activity may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of hematopoiesis is beneficial. For example, a protein of the present invention may be useful in regulation of hematopoiesis and, consequently, in the treatment of myeloid or lymphoid cell deficiencies. Even marginal biological activity in support of colony forming cells or of factor-dependent cell lines indicates involvement in regulating hematopoiesis, e.g. in supporting the growth and proliferation of erythroid progenitor cells alone or in combination with other cytokines, thereby indicating utility, for example, in treating various anemias or for use in conjunction with irradiation/chemotherapy to stimulate the production of erythroid precursors and/or erythroid cells; in supporting the growth and proliferation of myeloid cells such as granulocytes and monocytes/macrophages (i.e., traditional CSF activity) useful, for example, in conjunction with chemotherapy to prevent or treat consequent myelo-suppression; in supporting the growth and proliferation of megakaryocytes and consequently of platelets thereby allowing prevention or treatment of various platelet disorders such as thrombocytopenia, and generally for use in place of or complimentary to platelet transfusions; and/or in supporting the growth and proliferation of hematopoietic stem cells which are capable of maturing to any and all of the above-mentioned hematopoietic cells and therefore find therapeutic utility in various stem cell disorders (such as those usually treated with transplantation, including, without limitation, aplastic anemia and paroxysmal nocturnal hemoglobinuria), as well as in repopulating the stem cell compartment post irradiation/chemotherapy, either in-vivo or ex-vivo (i.e., in conjunction with bone marrow transplantation or with peripheral progenitor cell transplantation (homologous or heterologous)) as normal cells or genetically manipulated for gene therapy. Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

#### EXAMPLE 35

Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for  
Regulation of Tissue Growth

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their effect on tissue growth. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in International Patent Publication No. WO95/16035, International Patent Publication No. WO95/05846 and International Patent Publication No. WO91/07491, which are incorporated herein by reference.

Assays for wound healing activity include, without limitation, those described in: Winter, Epidermal Wound Healing, pps. 71-112 (Maibach, H1 and Rovee, DT, eds.), Year Book Medical Publishers, Inc., Chicago, as modified by Eaglstein and Mertz, J. Invest. Dermatol 71:382-84 (1978) which are incorporated herein by reference.

Those proteins which are involved in the regulation of tissue growth may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of tissue growth is beneficial. For example, a protein of the present invention also may have utility in compositions used for bone, cartilage, tendon, ligament and/or nerve tissue growth or regeneration, as well as for wound healing and tissue repair and replacement, and in the treatment of burns, incisions and ulcers.

A protein of the present invention, which induces cartilage and/or bone growth in circumstances where bone is not normally formed, has application in the healing of bone fractures and cartilage damage or defects in humans and other animals. Such a preparation employing a protein of the invention may have prophylactic use in closed as well as open fracture reduction and also in the improved fixation of artificial joints. De novo bone formation induced by an osteogenic agent contributes to the repair of congenital, trauma induced, or oncologic resection induced craniofacial defects, and also is useful in cosmetic plastic surgery.

A protein of this invention may also be used in the treatment of periodontal disease, and in other tooth repair processes. Such agents may provide an environment to attract bone-forming cells, stimulate growth of bone-forming cells or induce differentiation of progenitors of bone-forming cells. A protein of the invention may also

be useful in the treatment of osteoporosis or osteoarthritis, such as through stimulation of bone and/or cartilage repair or by blocking inflammation or processes of tissue destruction (collagenase activity, osteoclast activity, etc.) mediated by inflammatory processes.

Another category of tissue regeneration activity that may be attributable to the protein of the present invention is tendon/ligament formation. A protein of the present invention, which induces tendon/ligament-like tissue or other tissue formation in circumstances where such tissue is not normally formed, has application in the healing of tendon or ligament tears, deformities and other tendon or ligament defects in humans and other animals. Such a preparation employing a tendon/ligament-like tissue inducing protein may have prophylactic use in preventing damage to tendon or ligament tissue, as well as use in the improved fixation of tendon or ligament to bone or other tissues, and in repairing defects to tendon or ligament tissue. De novo tendon/ligament-like tissue formation induced by a composition of the present invention contributes to the repair of congenital, trauma induced, or other tendon or ligament defects of other origin, and is also useful in cosmetic plastic surgery for attachment or repair of tendons or ligaments. The compositions of the present invention may provide an environment to attract tendon- or ligament-forming cells, stimulate growth of tendon- or ligament-forming cells, induce differentiation of progenitors of tendon- or ligament-forming cells, or induce growth of tendon/ligament cells or progenitors ex vivo for return in vivo to effect tissue repair. The compositions of the invention may also be useful in the treatment of tendinitis, carpal tunnel syndrome and other tendon or ligament defects. The compositions may also include an appropriate matrix and/or sequestering agent as a carrier as is well known in the art.

The protein of the present invention may also be useful for proliferation of neural cells and for regeneration of nerve and brain tissue, i.e., for the treatment of central and peripheral nervous system diseases and neuropathies, as well as mechanical and traumatic disorders, which involve degeneration, death or trauma to neural cells or nerve tissue. More specifically, a protein may be used in the treatment of diseases of the peripheral nervous system, such as peripheral nerve injuries, peripheral neuropathy and localized

neuropathies, and central nervous system diseases, such as Alzheimer's, Parkinson's disease, Huntington's disease, amyotrophic lateral sclerosis, and Shy-Drager syndrome. Further conditions which may be treated in accordance with the present invention include mechanical and traumatic disorders, such as spinal cord disorders, head trauma and cerebrovascular diseases such as stroke. Peripheral neuropathies resulting from chemotherapy or other medical therapies may also be treatable using a protein of the invention.

Proteins of the invention may also be useful to promote better or faster closure of non-healing wounds, including without limitation pressure ulcers, ulcers associated with vascular insufficiency, surgical and traumatic wounds, and the like.

It is expected that a protein of the present invention may also exhibit activity for generation or regeneration of other tissues, such as organs (including, for example, pancreas, liver, intestine, kidney, skin, endothelium) muscle (smooth, skeletal or cardiac) and vascular (including vascular endothelium) tissue, or for promoting the growth of cells comprising such tissues. Part of the desired effects may be by inhibition or modulation of fibrotic scarring to allow normal tissue to generate. A protein of the invention may also exhibit angiogenic activity.

A protein of the present invention may also be useful for gut protection or regeneration and treatment of lung or liver fibrosis, reperfusion injury in various tissues, and conditions resulting from systemic cytokine damage.

A protein of the present invention may also be useful for promoting or inhibiting differentiation of tissues described above from precursor tissues or cells; or for inhibiting the growth of tissues described above.

Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

### EXAMPLE 36

#### Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Regulation of Reproductive Hormones or Cell Movement



The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their ability to regulate reproductive hormones, such as follicle stimulating hormone. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein  
5 by reference: Vale et al., **Endocrinology** 91:562-572, 1972; Ling et al., **Nature** 321:779-782, 1986; Vale et al., **Nature** 321:776-779, 1986; Mason et al., **Nature** 318:659-663, 1985; Forage et al., **Proc. Natl. Acad. Sci. USA** 83:3091-3095, 1986. Chapter 6.12 (Measurement of Alpha and Beta Chemokines) **Current Protocols in Immunology**, J.E. Coligan et al. Eds. Greene Publishing Associates and Wiley-Interscience; Taub et al. **J.**  
10 **Clin. Invest.** 95:1370-1376, 1995; Lind et al. **APMIS** 103:140-146, 1995; Muller et al. **Eur. J. Immunol.** 25:1744-1748; Gruber et al. **J. of Immunol.** 152:5860-5867, 1994; Johnston et al. **J. of Immunol.** 153:1762-1768, 1994.

Those proteins which exhibit activity as reproductive hormones or regulators of cell movement may then be formulated as pharmaceuticals and used to treat clinical  
15 conditions in which regulation of reproductive hormones or cell movement are beneficial. For example, a protein of the present invention may also exhibit activin- or inhibin-related activities. Inhibins are characterized by their ability to inhibit the release of follicle stimulating hormone (FSH), while activins are characterized by their ability to stimulate the release of follicle stimulating hormone (FSH). Thus, a protein of the present invention,  
20 alone or in heterodimers with a member of the inhibin  $\alpha$  family, may be useful as a contraceptive based on the ability of inhibins to decrease fertility in female mammals and decrease spermatogenesis in male mammals. Administration of sufficient amounts of other inhibins can induce infertility in these mammals. Alternatively, the protein of the invention, as a homodimer or as a heterodimer with other protein subunits of the inhibin-B  
25 group, may be useful as a fertility inducing therapeutic, based upon the ability of activin molecules in stimulating FSH release from cells of the anterior pituitary. See, for example, United States Patent 4,798,885, the disclosure of which is incorporated herein by reference. A protein of the invention may also be useful for advancement of the onset of

fertility in sexually immature mammals, so as to increase the lifetime reproductive performance of domestic animals such as cows, sheep and pigs.

Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

### EXAMPLE 36A

#### Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Chemotactic/Chemokinetic Activity

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for chemotactic/chemokinetic activity. For example, a protein of the present invention may have chemotactic or chemokinetic activity (e.g., act as a chemokine) for mammalian cells, including, for example, monocytes, fibroblasts, neutrophils, T-cells, mast cells, eosinophils, epithelial and/or endothelial cells. Chemotactic and chemokinetic proteins can be used to mobilize or attract a desired cell population to a desired site of action.

Chemotactic or chemokinetic proteins provide particular advantages in treatment of wounds and other trauma to tissues, as well as in treatment of localized infections. For example, attraction of lymphocytes, monocytes or neutrophils to tumors or sites of infection may result in improved immune responses against the tumor or infecting agent.

A protein or peptide has chemotactic activity for a particular cell population if it can stimulate, directly or indirectly, the directed orientation or movement of such cell population. Preferably, the protein or peptide has the ability to directly stimulate directed movement of cells. Whether a particular protein has chemotactic activity for a population of cells can be readily determined by employing such protein or peptide in any known assay for cell chemotaxis.

The activity of a protein of the invention may, among other means, be measured by the following methods:

Assays for chemotactic activity (which will identify proteins that induce or prevent chemotaxis) consist of assays that measure the ability of a protein to induce the migration of cells across a membrane as well as the ability of a protein to induce the adhesion of

one cell population to another cell population. Suitable assays for movement and adhesion include, without limitation, those described in: Current Protocols in Immunology, Ed by J.E. Coligan, A.M. Kruisbeek, D.H. Margulies, E.M. Shevach, W. Strober, Pub. Greene Publishing Associates and Wiley-Interscience (Chapter 6.12, Measurement of alpha and beta Chemokines 6.12.1-6.12.28; Taub et al. J. Clin. Invest. 95:1370-1376, 1995; Lind et al. APMIS 103:140-146, 1995; Mueller et al Eur. J. Immunol. 25:1744-1748; Gruber et al. J. of Immunol. 152:5860-5867, 1994; Johnston et al. J. of Immunol, 153:1762-1768, 1994.

### EXAMPLE 37

#### Assaying the Proteins Expressed from Extended cDNAs or Portions Thereof for Regulation of Blood Clotting

The proteins encoded by the extended cDNAs or portions thereof may also be evaluated for their effects on blood clotting. Numerous assays for such activity are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Linet et al., **J. Clin. Pharmacol.** 26:131-140, 1986; Burdick et al., **Thrombosis Res.** 45:413-419, 1987; Humphrey et al., **Fibrinolysis** 5:71-79 (1991); Schaub, **Prostaglandins** 35:467-474, 1988.

Those proteins which are involved in the regulation of blood clotting may then be formulated as pharmaceuticals and used to treat clinical conditions in which regulation of blood clotting is beneficial. For example, a protein of the invention may also exhibit hemostatic or thrombolytic activity. As a result, such a protein is expected to be useful in treatment of various coagulations disorders (including hereditary disorders, such as hemophilias) or to enhance coagulation and other hemostatic events in treating wounds resulting from trauma, surgery or other causes. A protein of the invention may also be useful for dissolving or inhibiting formation of thromboses and for treatment and prevention of conditions resulting therefrom (such as, for example, infarction of cardiac and central nervous system vessels (e.g., stroke). Alternatively, as described in more detail below, genes encoding these proteins or nucleic acids regulating the expression of

these proteins may be introduced into appropriate host cells to increase or decrease the expression of the proteins as desired.

### EXAMPLE 38

#### Assaying the Proteins Expressed from Extended cDNAs or

#### Portions Thereof for Involvement in Receptor/Ligand Interactions

5 The proteins encoded by the extended cDNAs or a portion thereof may also be evaluated for their involvement in receptor/ligand interactions. Numerous assays for such involvement are familiar to those skilled in the art, including the assays disclosed in the following references, which are incorporated herein by reference: Chapter 7.28  
10 (Measurement of Cellular Adhesion under Static Conditions 7.28.1-7.28.22) in **Current Protocols in Immunology**, J.E. Coligan et al. Eds. Greene Publishing Associates and Wiley-Interscience; Takai et al., **Proc. Natl. Acad. Sci. USA** 84:6864-6868, 1987; Bierer et al., **J. Exp. Med.** 168:1145-1156, 1988; Rosenstein et al., **J. Exp. Med.** 169:149-160, 1989; Stoltenborg et al., **J. Immunol. Methods** 175:59-68, 1994; Stitt et al., **Cell** 80:661-  
15 670, 1995; Gyuris et al., **Cell** 75:791-803, 1993.

For example, the proteins of the present invention may also demonstrate activity as receptors, receptor ligands or inhibitors or agonists of receptor/ligand interactions. Examples of such receptors and ligands include, without limitation, cytokine receptors and their ligands, receptor kinases and their ligands, receptor phosphatases and their ligands,  
20 receptors involved in cell-cell interactions and their ligands (including without limitation, cellular adhesion molecules (such as selectins, integrins and their ligands) and receptor/ligand pairs involved in antigen presentation, antigen recognition and development of cellular and humoral immune responses). Receptors and ligands are also useful for screening of potential peptide or small molecule inhibitors of the relevant  
25 receptor/ligand interaction. A protein of the present invention (including, without limitation, fragments of receptors and ligands) may themselves be useful as inhibitors of receptor/ligand interactions.

### EXAMPLE 38A

#### Assaying the Proteins Expressed from Extended cDNAs or Portions

### Thereof for Anti-Inflammatory Activity

The proteins encoded by the extended cDNAs or a portion thereof may also be evaluated for anti-inflammatory activity. The anti-inflammatory activity may be achieved by providing a stimulus to cells involved in the inflammatory response, by inhibiting or promoting cell-cell interactions (such as, for example, cell adhesion), by inhibiting or promoting chemotaxis of cells involved in the inflammatory process, inhibiting or promoting cell extravasation, or by stimulating or suppressing production of other factors which more directly inhibit or promote an inflammatory response. Proteins exhibiting such activities can be used to treat inflammatory conditions including chronic or acute conditions), including without limitation inflammation associated with infection (such as septic shock, sepsis or systemic inflammatory response syndrome (SIRS)), ischemia-reperfusion injury, endotoxin lethality, arthritis, complement-mediated hyperacute rejection, nephritis, cytokine or chemokine-induced lung injury, inflammatory bowel disease, Crohn's disease or resulting from over production of cytokines such as TNF or IL-1. Proteins of the invention may also be useful to treat anaphylaxis and hypersensitivity to an antigenic substance or material.

### **EXAMPLE 38B**

#### Assaying the Proteins Expressed from Extended cDNAs or

#### Portions Thereof for Tumor Inhibition Activity

The proteins encoded by the extended cDNAs or a portion thereof may also be evaluated for tumor inhibition activity. In addition to the activities described above for immunological treatment or prevention of tumors, a protein of the invention may exhibit other anti-tumor activities. A protein may inhibit tumor growth directly or indirectly (such as, for example, via ADCC). A protein may exhibit its tumor inhibitory activity by acting on tumor tissue or tumor precursor tissue, by inhibiting formation of tissues necessary to support tumor growth (such as, for example, by inhibiting angiogenesis), by causing production of other factors, agents or cell types which inhibit tumor growth, or by suppressing, eliminating or inhibiting factors, agents or cell types which promote tumor growth.

A protein of the invention may also exhibit one or more of the following additional activities or effects: inhibiting the growth, infection or function of, or killing, infectious agents, including, without limitation, bacteria, viruses, fungi and other parasites; effecting (suppressing or enhancing) bodily characteristics, including, without limitation, height, weight, hair color, eye color, skin, fat to lean ratio or other tissue pigmentation, or organ or body part size or shape (such as, for example, breast augmentation or diminution, change in bone form or shape); effecting biorhythms or circadian cycles or rhythms; effecting the fertility of male or female subjects; effecting the metabolism, catabolism, anabolism, processing, utilization, storage or elimination of dietary fat, lipid, protein, carbohydrate, vitamins, minerals, cofactors or other nutritional factors or component(s); effecting behavioral characteristics, including, without limitation, appetite, libido, stress, cognition (including cognitive disorders), depression (including depressive disorders) and violent behaviors; providing analgesic effects or other pain reducing effects; promoting differentiation and growth of embryonic stem cells in lineages other than hematopoietic lineages; hormonal or endocrine activity; in the case of enzymes, correcting deficiencies of the enzyme and treating deficiency-related diseases; treatment of hyperproliferative disorders (such as, for example, psoriasis); immunoglobulin-like activity (such as, for example, the ability to bind antigens or complement); and the ability to act as an antigen in a vaccine composition to raise an immune response against such protein or another material or entity which is cross-reactive with such protein.

### **EXAMPLE 39**

#### **Identification of Proteins which Interact with Polypeptides Encoded by Extended cDNAs**

Proteins which interact with the polypeptides encoded by extended cDNAs or portions thereof, such as receptor proteins, may be identified using two hybrid systems such as the Matchmaker Two Hybrid System 2 (Catalog No. K1604-1, Clontech). As described in the manual accompanying the Matchmaker Two Hybrid System 2 (Catalog No. K1604-1, Clontech), which is incorporated herein by reference, the extended cDNAs or portions thereof, are inserted into an expression vector such that they are in frame with

DNA encoding the DNA binding domain of the yeast transcriptional activator GAL4. cDNAs in a cDNA library which encode proteins which might interact with the polypeptides encoded by the extended cDNAs or portions thereof are inserted into a second expression vector such that they are in frame with DNA encoding the activation domain of GAL4. The two expression plasmids are transformed into yeast and the yeast are plated on selection medium which selects for expression of selectable markers on each of the expression vectors as well as GAL4 dependent expression of the HIS3 gene. Transformants capable of growing on medium lacking histidine are screened for GAL4 dependent lacZ expression. Those cells which are positive in both the histidine selection and the lacZ assay contain plasmids encoding proteins which interact with the polypeptide encoded by the extended cDNAs or portions thereof.

Alternatively, the system described in Lustig et al., Methods in Enzymology 283: 83-99 (1997), the disclosure of which is incorporated herein by reference, may be used for identifying molecules which interact with the polypeptides encoded by extended cDNAs. In such systems, *in vitro* transcription reactions are performed on a pool of vectors containing extended cDNA inserts cloned downstream of a promoter which drives *in vitro* transcription. The resulting pools of mRNAs are introduced into *Xenopus laevis* oocytes. The oocytes are then assayed for a desired activity.

Alternatively, the pooled *in vitro* transcription products produced as described above may be translated *in vitro*. The pooled *in vitro* translation products can be assayed for a desired activity or for interaction with a known polypeptide.

Proteins or other molecules interacting with polypeptides encoded by extended cDNAs can be found by a variety of additional techniques. In one method, affinity columns containing the polypeptide encoded by the extended cDNA or a portion thereof can be constructed. In some versions, of this method the affinity column contains chimeric proteins in which the protein encoded by the extended cDNA or a portion thereof is fused to glutathione S-transferase. A mixture of cellular proteins or pool of expressed proteins as described above and is applied to the affinity column. Proteins interacting with the polypeptide attached to the column can then be isolated and

analyzed on 2-D electrophoresis gel as described in Ramunsen et al. Electrophoresis, 18, 588-598 (1997), the disclosure of which is incorporated herein by reference. Alternatively, the proteins retained on the affinity column can be purified by electrophoresis based methods and sequenced. The same method can be used to isolate antibodies, to screen phage display products, or to screen phage display human antibodies.

Proteins interacting with polypeptides encoded by extended cDNAs or portions thereof can also be screened by using an Optical Biosensor as described in Edwards & Leatherbarrow, Analytical Biochemistry, 246, 1-6 (1997), the disclosure of which is incorporated herein by reference. The main advantage of the method is that it allows the determination of the association rate between the protein and other interacting molecules. Thus, it is possible to specifically select interacting molecules with a high or low association rate. Typically a target molecule is linked to the sensor surface (through a carboxymethyl dextran matrix) and a sample of test molecules is placed in contact with the target molecules. The binding of a test molecule to the target molecule causes a change in the refractive index and/ or thickness. This change is detected by the Biosensor provided it occurs in the evanescent field (which extend a few hundred nanometers from the sensor surface). In these screening assays, the target molecule can be one of the polypeptides encoded by extended cDNAs or a portion thereof and the test sample can be a collection of proteins extracted from tissues or cells, a pool of expressed proteins, combinatorial peptide and/ or chemical libraries, or phage displayed peptides. The tissues or cells from which the test proteins are extracted can originate from any species.

In other methods, a target protein is immobilized and the test population is a collection of unique polypeptides encoded by the extended cDNAs or portions thereof.

To study the interaction of the proteins encoded by the extended cDNAs or portions thereof with drugs, the microdialysis coupled to HPLC method described by Wang et al., Chromatographia, 44, 205-208(1997) or the affinity capillary



electrophoresis method described by Busch et al., J. Chromatogr. 777:311-328 (1997), the disclosures of which are incorporated herein by reference can be used.

5 The system described in U.S. Patent No. 5,654,150, the disclosure of which is incorporated herein by reference, may also be used to identify molecules which interact with the polypeptides encoded by the extended cDNAs. In this system, pools of extended cDNAs are transcribed and translated *in vitro* and the reaction products are assayed for interaction with a known polypeptide or antibody.

10 It will be appreciated by those skilled in the art that the proteins expressed from the extended cDNAs or portions may be assayed for numerous activities in addition to those specifically enumerated above. For example, the expressed proteins may be evaluated for applications involving control and regulation of inflammation, tumor proliferation or metastasis, infection, or other clinical conditions. In addition, the proteins expressed from the extended cDNAs or portions thereof may be useful as nutritional agents or cosmetic agents.

15 The proteins expressed from the extended cDNAs or portions thereof may be used to generate antibodies capable of specifically binding to the expressed protein or fragments thereof as described in Example 40 below. The antibodies may be capable of binding a full length protein encoded by one of the sequences of SEQ ID NOs. 40-140 and 242-377, a mature protein encoded by one of the sequences of SEQ ID NOs. 40-140 and 20 242-377, or a signal peptide encoded by one of the sequences of SEQ ID Nos. 40-140 and 242-377. Alternatively, the antibodies may be capable of binding fragments of the proteins expressed from the extended cDNAs which comprise at least 10 amino acids of the sequences of SEQ ID NOs: 141-241 and 378-513. In some embodiments, the antibodies may be capable of binding fragments of the proteins expressed from the 25 extended cDNAs which comprise at least 15 amino acids of the sequences of SEQ ID NOs: 141-241 and 378-513. In other embodiments, the antibodies may be capable of binding fragments of the proteins expressed from the extended cDNAs which comprise at least 25 amino acids of the sequences of SEQ ID NOs: 141-241 and 378-513. In further embodiments, the antibodies may be capable of binding fragments of the proteins

expressed from the extended cDNAs which comprise at least 40 amino acids of the sequences of SEQ ID NOs: 141-241 and 378-513.

#### EXAMPLE 40

##### Production of an Antibody to a Human Protein

5 Substantially pure protein or polypeptide is isolated from the transfected or transformed cells as described in Example 30. The concentration of protein in the final preparation is adjusted, for example, by concentration on an Amicon filter device, to the level of a few micrograms/ml. Monoclonal or polyclonal antibody to the protein can then be prepared as follows:

##### 10 A. Monoclonal Antibody Production by Hybridoma Fusion

Monoclonal antibody to epitopes of any of the peptides identified and isolated as described can be prepared from murine hybridomas according to the classical method of Kohler, G. and Milstein, C., **Nature** 256:495 (1975) or derivative methods thereof. Briefly, a mouse is repetitively inoculated with a few micrograms of the selected protein or peptides derived therefrom over a period of a few weeks. The mouse is then sacrificed, and the antibody producing cells of the spleen isolated. The spleen cells are fused by means of polyethylene glycol with mouse myeloma cells, and the excess unfused cells destroyed by growth of the system on selective media comprising aminopterin (HAT media). The successfully fused cells are diluted and aliquots of the dilution placed in wells of a microtiter plate where growth of the culture is continued. Antibody-producing clones are identified by detection of antibody in the supernatant fluid of the wells by immunoassay procedures, such as Elisa, as originally described by Engvall, E., **Meth. Enzymol.** 70:419 (1980), and derivative methods thereof. Selected positive clones can be expanded and their monoclonal antibody product harvested for use. Detailed procedures for monoclonal antibody production are described in Davis, L. et al. **Basic Methods in Molecular Biology** Elsevier, New York. Section 21-2.

##### 25 B. Polyclonal Antibody Production by Immunization

Polyclonal antiserum containing antibodies to heterogenous epitopes of a single protein can be prepared by immunizing suitable animals with the expressed protein or

peptides derived therefrom described above, which can be unmodified or modified to enhance immunogenicity. Effective polyclonal antibody production is affected by many factors related both to the antigen and the host species. For example, small molecules tend to be less immunogenic than others and may require the use of carriers and adjuvant.

5 Also, host animals vary in response to site of inoculations and dose, with both inadequate or excessive doses of antigen resulting in low titer antisera. Small doses (ng level) of antigen administered at multiple intradermal sites appears to be most reliable. An effective immunization protocol for rabbits can be found in Vaitukaitis, J. et al. **J. Clin. Endocrinol. Metab.** **33**:988-991 (1971).

10           Booster injections can be given at regular intervals, and antiserum harvested when antibody titer thereof, as determined semi-quantitatively, for example, by double immunodiffusion in agar against known concentrations of the antigen, begins to fall. See, for example, Ouchterlony, O. et al., Chap. 19 in: **Handbook of Experimental Immunology** D. Wier (ed) Blackwell (1973). Plateau concentration of antibody is usually  
15 in the range of 0.1 to 0.2 mg/ml of serum (about 12  $\mu$ M). Affinity of the antisera for the antigen is determined by preparing competitive binding curves, as described, for example, by Fisher, D., Chap. 42 in: **Manual of Clinical Immunology**, 2d Ed. (Rose and Friedman, Eds.) Amer. Soc. For Microbiol., Washington, D.C. (1980).

20           Antibody preparations prepared according to either protocol are useful in quantitative immunoassays which determine concentrations of antigen-bearing substances in biological samples; they are also used semi-quantitatively or qualitatively to identify the presence of antigen in a biological sample. The antibodies may also be used in therapeutic compositions for killing cells expressing the protein or reducing the levels of the protein in the body.

## 25   **V. Use of Extended cDNAs or Portions Thereof as Reagents**

          The extended cDNAs of the present invention may be used as reagents in isolation procedures, diagnostic assays, and forensic procedures. For example, sequences from the extended cDNAs (or genomic DNAs obtainable therefrom) may be detectably labeled and used as probes to isolate other sequences capable of hybridizing to them. In addition,

sequences from the extended cDNAs (or genomic DNAs obtainable therefrom) may be used to design PCR primers to be used in isolation, diagnostic, or forensic procedures.

#### EXAMPLE 41

##### Preparation of PCR Primers and Amplification of DNA

5           The extended cDNAs (or genomic DNAs obtainable therefrom) may be used to prepare PCR primers for a variety of applications, including isolation procedures for cloning nucleic acids capable of hybridizing to such sequences, diagnostic techniques and forensic techniques. The PCR primers are at least 10 bases, and preferably at least 12, 15, or 17 bases in length. More preferably, the PCR primers are at least 20-30 bases in length.

10          In some embodiments, the PCR primers may be more than 30 bases in length. It is preferred that the primer pairs have approximately the same G/C ratio, so that melting temperatures are approximately the same. A variety of PCR techniques are familiar to those skilled in the art. For a review of PCR technology, see Molecular Cloning to Genetic Engineering White, B.A. Ed. in Methods in Molecular Biology 67: Humana Press, Totowa 1997. In each of these PCR procedures, PCR primers on either side of the  
15          nucleic acid sequences to be amplified are added to a suitably prepared nucleic acid sample along with dNTPs and a thermostable polymerase such as Taq polymerase, Pfu polymerase, or Vent polymerase. The nucleic acid in the sample is denatured and the PCR primers are specifically hybridized to complementary nucleic acid sequences in the  
20          sample. The hybridized primers are extended. Thereafter, another cycle of denaturation, hybridization, and extension is initiated. The cycles are repeated multiple times to produce an amplified fragment containing the nucleic acid sequence between the primer sites.

#### EXAMPLE 42

##### Use of Extended cDNAs as Probes

25           Probes derived from extended cDNAs or portions thereof (or genomic DNAs obtainable therefrom) may be labeled with detectable labels familiar to those skilled in the art, including radioisotopes and non-radioactive labels, to provide a detectable probe. The detectable probe may be single stranded or double stranded and may be made using

techniques known in the art, including in vitro transcription, nick translation, or kinase reactions. A nucleic acid sample containing a sequence capable of hybridizing to the labeled probe is contacted with the labeled probe. If the nucleic acid in the sample is double stranded, it may be denatured prior to contacting the probe. In some applications, the nucleic acid sample may be immobilized on a surface such as a nitrocellulose or nylon membrane. The nucleic acid sample may comprise nucleic acids obtained from a variety of sources, including genomic DNA, cDNA libraries, RNA, or tissue samples.

Procedures used to detect the presence of nucleic acids capable of hybridizing to the detectable probe include well known techniques such as Southern blotting, Northern blotting, dot blotting, colony hybridization, and plaque hybridization. In some applications, the nucleic acid capable of hybridizing to the labeled probe may be cloned into vectors such as expression vectors, sequencing vectors, or in vitro transcription vectors to facilitate the characterization and expression of the hybridizing nucleic acids in the sample. For example, such techniques may be used to isolate and clone sequences in a genomic library or cDNA library which are capable of hybridizing to the detectable probe as described in Example 30 above.

PCR primers made as described in Example 41 above may be used in forensic analyses, such as the DNA fingerprinting techniques described in Examples 43-47 below. Such analyses may utilize detectable probes or primers based on the sequences of the extended cDNAs isolated using the 5' ESTs (or genomic DNAs obtainable therefrom).

### **EXAMPLE 43**

#### **Forensic Matching by DNA Sequencing**

In one exemplary method, DNA samples are isolated from forensic specimens of, for example, hair, semen, blood or skin cells by conventional methods. A panel of PCR primers based on a number of the extended cDNAs (or genomic DNAs obtainable therefrom), is then utilized in accordance with Example 41 to amplify DNA of approximately 100-200 bases in length from the forensic specimen. Corresponding sequences are obtained from a test subject. Each of these identification DNAs is then sequenced using standard techniques, and a simple database comparison determines the

differences, if any, between the sequences from the subject and those from the sample. Statistically significant differences between the suspect's DNA sequences and those from the sample conclusively prove a lack of identity. This lack of identity can be proven, for example, with only one sequence. Identity, on the other hand, should be demonstrated with a large number of sequences, all matching. Preferably, a minimum of 50 statistically identical sequences of 100 bases in length are used to prove identity between the suspect and the sample.

#### **EXAMPLE 44**

##### Positive Identification by DNA Sequencing

The technique outlined in the previous example may also be used on a larger scale to provide a unique fingerprint-type identification of any individual. In this technique, primers are prepared from a large number of sequences from Table IV and the appended sequence listing. Preferably, 20 to 50 different primers are used. These primers are used to obtain a corresponding number of PCR-generated DNA segments from the individual in question in accordance with Example 41. Each of these DNA segments is sequenced, using the methods set forth in Example 43. The database of sequences generated through this procedure uniquely identifies the individual from whom the sequences were obtained. The same panel of primers may then be used at any later time to absolutely correlate tissue or other biological specimen with that individual.

#### **EXAMPLE 45**

##### Southern Blot Forensic Identification

The procedure of Example 44 is repeated to obtain a panel of at least 10 amplified sequences from an individual and a specimen. Preferably, the panel contains at least 50 amplified sequences. More preferably, the panel contains 100 amplified sequences. In some embodiments, the panel contains 200 amplified sequences. This PCR-generated DNA is then digested with one or a combination of, preferably, four base specific restriction enzymes. Such enzymes are commercially available and known to those of skill in the art. After digestion, the resultant gene fragments are size separated in multiple duplicate wells on an agarose gel and transferred to nitrocellulose using Southern blotting

techniques well known to those with skill in the art. For a review of Southern blotting see Davis et al. (**Basic Methods in Molecular Biology**, 1986, Elsevier Press. pp 62-65).

5 A panel of probes based on the sequences of the extended cDNAs (or genomic DNAs obtainable therefrom), or fragments thereof of at least 10 bases, are radioactively or colorimetrically labeled using methods known in the art, such as nick translation or end labeling, and hybridized to the Southern blot using techniques known in the art (Davis et al., supra). Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). More preferably, the probe comprises at least 20-30 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). In some embodiments, the probe comprises more than 30 nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). In other embodiments, the probe comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom).

15 Preferably, at least 5 to 10 of these labeled probes are used, and more preferably at least about 20 or 30 are used to provide a unique pattern. The resultant bands appearing from the hybridization of a large sample of extended cDNAs (or genomic DNAs obtainable therefrom) will be a unique identifier. Since the restriction enzyme cleavage will be different for every individual, the band pattern on the Southern blot will also be unique. Increasing the number of extended cDNA probes will provide a statistically higher level of confidence in the identification since there will be an increased number of sets of bands used for identification.

#### **EXAMPLE 46**

##### **Dot Blot Identification Procedure**

25 Another technique for identifying individuals using the extended cDNA sequences disclosed herein utilizes a dot blot hybridization technique.

Genomic DNA is isolated from nuclei of subject to be identified. Oligonucleotide probes of approximately 30 bp in length are synthesized that correspond to at least 10, preferably 50 sequences from the extended cDNAs or genomic DNAs obtainable

therefrom. The probes are used to hybridize to the genomic DNA through conditions known to those in the art. The oligonucleotides are end labeled with  $P^{32}$  using polynucleotide kinase (Pharmacia). Dot Blots are created by spotting the genomic DNA onto nitrocellulose or the like using a vacuum dot blot manifold (BioRad, Richmond California). The nitrocellulose filter containing the genomic sequences is baked or UV linked to the filter, prehybridized and hybridized with labeled probe using techniques known in the art (Davis et al. supra). The  $^{32}P$  labeled DNA fragments are sequentially hybridized with successively stringent conditions to detect minimal differences between the 30 bp sequence and the DNA. Tetramethylammonium chloride is useful for identifying clones containing small numbers of nucleotide mismatches (Wood et al., **Proc. Natl. Acad. Sci. USA** **82(6)**:1585-1588 (1985)) which is hereby incorporated by reference. A unique pattern of dots distinguishes one individual from another individual.

Extended cDNAs or oligonucleotides containing at least 10 consecutive bases from these sequences can be used as probes in the following alternative fingerprinting technique. Preferably, the probe comprises at least 12, 15, or 17 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). More preferably, the probe comprises at least 20-30 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). In some embodiments, the probe comprises more than 30 nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom). In other embodiments, the probe comprises at least 40, at least 50, at least 75, at least 100, at least 150, or at least 200 consecutive nucleotides from the extended cDNA (or genomic DNAs obtainable therefrom).

Preferably, a plurality of probes having sequences from different genes are used in the alternative fingerprinting technique. Example 47 below provides a representative alternative fingerprinting procedure in which the probes are derived from extended cDNAs.

#### **EXAMPLE 47**

##### Alternative "Fingerprint" Identification Technique



20-mer oligonucleotides are prepared from a large number, e.g. 50, 100, or 200, of extended cDNA sequences (or genomic DNAs obtainable therefrom) using commercially available oligonucleotide services such as Genset, Paris, France. Cell samples from the test subject are processed for DNA using techniques well known to those with skill in the art. The nucleic acid is digested with restriction enzymes such as EcoRI and XbaI. Following digestion, samples are applied to wells for electrophoresis. The procedure, as known in the art, may be modified to accommodate polyacrylamide electrophoresis, however in this example, samples containing 5 ug of DNA are loaded into wells and separated on 0.8% agarose gels. The gels are transferred onto nitrocellulose using standard Southern blotting techniques.

10 ng of each of the oligonucleotides are pooled and end-labeled with P<sup>32</sup>. The nitrocellulose is prehybridized with blocking solution and hybridized with the labeled probes. Following hybridization and washing, the nitrocellulose filter is exposed to X-Omat AR X-ray film. The resulting hybridization pattern will be unique for each individual.

It is additionally contemplated within this example that the number of probe sequences used can be varied for additional accuracy or clarity.

The antibodies generated in Examples 30 and 40 above may be used to identify the tissue type or cell species from which a sample is derived as described above.

#### **EXAMPLE 48**

##### **Identification of Tissue Types or Cell Species by Means of Labeled Tissue Specific Antibodies**

Identification of specific tissues is accomplished by the visualization of tissue specific antigens by means of antibody preparations according to Examples 30 and 40 which are conjugated, directly or indirectly to a detectable marker. Selected labeled antibody species bind to their specific antigen binding partner in tissue sections, cell suspensions, or in extracts of soluble proteins from a tissue sample to provide a pattern for qualitative or semi-qualitative interpretation.

Antisera for these procedures must have a potency exceeding that of the native preparation, and for that reason, antibodies are concentrated to a mg/ml level by isolation of the gamma globulin fraction, for example, by ion-exchange chromatography or by ammonium sulfate fractionation. Also, to provide the most specific antisera, unwanted antibodies, for example to common proteins, must be removed from the gamma globulin fraction, for example by means of insoluble immunoabsorbents, before the antibodies are labeled with the marker. Either monoclonal or heterologous antisera is suitable for either procedure.

#### A. Immunohistochemical Techniques

Purified, high-titer antibodies, prepared as described above, are conjugated to a detectable marker, as described, for example, by Fudenberg, H., Chap. 26 in: **Basic Clinical Immunology**, 3rd Ed. Lange, Los Altos, California (1980) or Rose, N. et al., Chap. 12 in: **Methods in Immunodiagnosis**, 2d Ed. John Wiley 503 Sons, New York (1980).

A fluorescent marker, either fluorescein or rhodamine, is preferred, but antibodies can also be labeled with an enzyme that supports a color producing reaction with a substrate, such as horseradish peroxidase. Markers can be added to tissue-bound antibody in a second step, as described below. Alternatively, the specific antitissue antibodies can be labeled with ferritin or other electron dense particles, and localization of the ferritin coupled antigen-antibody complexes achieved by means of an electron microscope. In yet another approach, the antibodies are radiolabeled, with, for example  $^{125}\text{I}$ , and detected by overlaying the antibody treated preparation with photographic emulsion.

Preparations to carry out the procedures can comprise monoclonal or polyclonal antibodies to a single protein or peptide identified as specific to a tissue type, for example, brain tissue, or antibody preparations to several antigenically distinct tissue specific antigens can be used in panels, independently or in mixtures, as required.

Tissue sections and cell suspensions are prepared for immunohistochemical examination according to common histological techniques. Multiple cryostat sections (about 4  $\mu\text{m}$ , unfixed) of the unknown tissue and known control, are mounted and each

slide covered with different dilutions of the antibody preparation. Sections of known and unknown tissues should also be treated with preparations to provide a positive control, a negative control, for example, pre-immune sera, and a control for non-specific staining, for example, buffer.

5           Treated sections are incubated in a humid chamber for 30 min at room temperature, rinsed, then washed in buffer for 30-45 min. Excess fluid is blotted away, and the marker developed.

          If the tissue specific antibody was not labeled in the first incubation, it can be labeled at this time in a second antibody-antibody reaction, for example, by adding  
10       fluorescein- or enzyme-conjugated antibody against the immunoglobulin class of the antiserum-producing species, for example, fluorescein labeled antibody to mouse IgG. Such labeled sera are commercially available.

          The antigen found in the tissues by the above procedure can be quantified by measuring the intensity of color or fluorescence on the tissue section, and calibrating that  
15       signal using appropriate standards.

#### **B.       Identification of Tissue Specific Soluble Proteins**

          The visualization of tissue specific proteins and identification of unknown tissues from that procedure is carried out using the labeled antibody reagents and detection strategy as described for immunohistochemistry; however the sample is prepared  
20       according to an electrophoretic technique to distribute the proteins extracted from the tissue in an orderly array on the basis of molecular weight for detection.

          A tissue sample is homogenized using a Virtis apparatus; cell suspensions are disrupted by Dounce homogenization or osmotic lysis, using detergents in either case as required to disrupt cell membranes, as is the practice in the art. Insoluble cell components  
25       such as nuclei, microsomes, and membrane fragments are removed by ultracentrifugation, and the soluble protein-containing fraction concentrated if necessary and reserved for analysis.

          A sample of the soluble protein solution is resolved into individual protein species by conventional SDS polyacrylamide electrophoresis as described, for example, by Davis,

L. et al., Section 19-2 in: **Basic Methods in Molecular Biology** (P. Leder, ed), Elsevier, New York (1986), using a range of amounts of polyacrylamide in a set of gels to resolve the entire molecular weight range of proteins to be detected in the sample. A size marker is run in parallel for purposes of estimating molecular weights of the constituent proteins.

5 Sample size for analysis is a convenient volume of from 5 to 55  $\mu$ l, and containing from about 1 to 100  $\mu$ g protein. An aliquot of each of the resolved proteins is transferred by blotting to a nitrocellulose filter paper, a process that maintains the pattern of resolution. Multiple copies are prepared. The procedure, known as Western Blot Analysis, is well described in Davis, L. et al., (above) Section 19-3. One set of nitrocellulose blots is  
10 stained with Coomassie Blue dye to visualize the entire set of proteins for comparison with the antibody bound proteins. The remaining nitrocellulose filters are then incubated with a solution of one or more specific antisera to tissue specific proteins prepared as described in Examples 30 and 40. In this procedure, as in procedure A above, appropriate positive and negative sample and reagent controls are run.

15 In either procedure A or B, a detectable label can be attached to the primary tissue antigen-primary antibody complex according to various strategies and permutations thereof. In a straightforward approach, the primary specific antibody can be labeled; alternatively, the unlabeled complex can be bound by a labeled secondary anti-IgG antibody. In other approaches, either the primary or secondary antibody is conjugated to a  
20 biotin molecule, which can, in a subsequent step, bind an avidin conjugated marker. According to yet another strategy, enzyme labeled or radioactive protein A, which has the property of binding to any IgG, is bound in a final step to either the primary or secondary antibody.

25 The visualization of tissue specific antigen binding at levels above those seen in control tissues to one or more tissue specific antibodies, prepared from the gene sequences identified from extended cDNA sequences, can identify tissues of unknown origin, for example, forensic samples, or differentiated tumor tissue that has metastasized to foreign bodily sites.

In addition to their applications in forensics and identification, extended cDNAs (or genomic DNAs obtainable therefrom) may be mapped to their chromosomal locations. Example 49 below describes radiation hybrid (RH) mapping of human chromosomal regions using extended cDNAs. Example 50 below describes a representative procedure for mapping an extended cDNA (or a genomic DNA obtainable therefrom) to its location on a human chromosome. Example 51 below describes mapping of extended cDNAs (or genomic DNAs obtainable therefrom) on metaphase chromosomes by Fluorescence In Situ Hybridization (FISH).

#### EXAMPLE 49

##### Radiation hybrid mapping of Extended cDNAs to the human genome

Radiation hybrid (RH) mapping is a somatic cell genetic approach that can be used for high resolution mapping of the human genome. In this approach, cell lines containing one or more human chromosomes are lethally irradiated, breaking each chromosome into fragments whose size depends on the radiation dose. These fragments are rescued by fusion with cultured rodent cells, yielding subclones containing different portions of the human genome. This technique is described by Benham et al. (*Genomics* 4:509-517, 1989) and Cox et al., (*Science* 250:245-250, 1990), the entire contents of which are hereby incorporated by reference. The random and independent nature of the subclones permits efficient mapping of any human genome marker. Human DNA isolated from a panel of 80-100 cell lines provides a mapping reagent for ordering extended cDNAs (or genomic DNAs obtainable therefrom). In this approach, the frequency of breakage between markers is used to measure distance, allowing construction of fine resolution maps as has been done using conventional ESTs (Schuler et al., *Science* 274:540-546, 1996, hereby incorporated by reference).

RH mapping has been used to generate a high-resolution whole genome radiation hybrid map of human chromosome 17q22-q25.3 across the genes for growth hormone (GH) and thymidine kinase (TK) (Foster et al., *Genomics* 33:185-192, 1996), the region surrounding the Gorlin syndrome gene (Obermayr et al., *Eur. J. Hum. Genet.* 4:242-245, 1996), 60 loci covering the entire short arm of chromosome 12 (Raeymaekers et al.,

*Genomics* 29:170-178, 1995), the region of human chromosome 22 containing the neurofibromatosis type 2 locus (Frazer et al., *Genomics* 14:574-584, 1992) and 13 loci on the long arm of chromosome 5 (Warrington et al., *Genomics* 11:701-708, 1991).

#### EXAMPLE 50

##### 5                                    Mapping of Extended cDNAs to Human    Chromosomes using PCR techniques

Extended cDNAs (or genomic DNAs obtainable therefrom) may be assigned to human chromosomes using PCR based methodologies. In such approaches, oligonucleotide primer pairs are designed from the extended cDNA sequence (or the  
10                                    sequence of a genomic DNA obtainable therefrom) to minimize the chance of amplifying through an intron. Preferably, the oligonucleotide primers are 18-23 bp in length and are designed for PCR amplification. The creation of PCR primers from known sequences is well known to those with skill in the art. For a review of PCR technology see Erlich, H.A., **PCR Technology; Principles and Applications for DNA Amplification**. 1992.  
15                                    W.H. Freeman and Co., New York.

The primers are used in polymerase chain reactions (PCR) to amplify templates from total human genomic DNA. PCR conditions are as follows: 60 ng of genomic DNA is used as a template for PCR with 80 ng of each oligonucleotide primer, 0.6 unit of Taq polymerase, and 1  $\mu$ Ci of a  $^{32}$ P-labeled deoxycytidine triphosphate. The PCR is  
20                                    performed in a microplate thermocycler (Techne) under the following conditions: 30 cycles of 94°C, 1.4 min; 55°C, 2 min; and 72°C, 2 min; with a final extension at 72°C for 10 min. The amplified products are analyzed on a 6% polyacrylamide sequencing gel and visualized by autoradiography. If the length of the resulting PCR product is identical to the distance between the ends of the primer sequences in the extended cDNA from which  
25                                    the primers are derived, then the PCR reaction is repeated with DNA templates from two panels of human-rodent somatic cell hybrids, BIOS PCRable DNA (BIOS Corporation) and NIGMS Human-Rodent Somatic Cell Hybrid Mapping Panel Number 1 (NIGMS, Camden, NJ).

PCR is used to screen a series of somatic cell hybrid cell lines containing defined sets of human chromosomes for the presence of a given extended cDNA (or genomic DNA obtainable therefrom). DNA is isolated from the somatic hybrids and used as starting templates for PCR reactions using the primer pairs from the extended cDNAs (or genomic DNAs obtainable therefrom). Only those somatic cell hybrids with chromosomes containing the human gene corresponding to the extended cDNA (or genomic DNA obtainable therefrom) will yield an amplified fragment. The extended cDNAs (or genomic DNAs obtainable therefrom) are assigned to a chromosome by analysis of the segregation pattern of PCR products from the somatic hybrid DNA templates. The single human chromosome present in all cell hybrids that give rise to an amplified fragment is the chromosome containing that extended cDNA (or genomic DNA obtainable therefrom). For a review of techniques and analysis of results from somatic cell gene mapping experiments. (See Ledbetter et al., **Genomics** 6:475-481 (1990).)

Alternatively, the extended cDNAs (or genomic DNAs obtainable therefrom) may be mapped to individual chromosomes using FISH as described in Example 51 below.

### EXAMPLE 51

#### Mapping of Extended 5' ESTs to Chromosomes

##### Using Fluorescence in situ Hybridization

Fluorescence in situ hybridization allows the extended cDNA (or genomic DNA obtainable therefrom) to be mapped to a particular location on a given chromosome. The chromosomes to be used for fluorescence in situ hybridization techniques may be obtained from a variety of sources including cell cultures, tissues, or whole blood.

In a preferred embodiment, chromosomal localization of an extended cDNA (or genomic DNA obtainable therefrom) is obtained by FISH as described by Cherif et al. (*Proc. Natl. Acad. Sci. U.S.A.*, **87**:6639-6643, 1990). Metaphase chromosomes are prepared from phytohemagglutinin (PHA)-stimulated blood cell donors. PHA-stimulated lymphocytes from healthy males are cultured for 72 h in RPMI-1640 medium. For synchronization, methotrexate (10  $\mu$ M) is added for 17 h, followed by addition of 5-bromodeoxyuridine (5-BudR, 0.1 mM) for 6 h. Colcemid (1  $\mu$ g/ml) is added for the last

15 min before harvesting the cells. Cells are collected, washed in RPMI, incubated with a hypotonic solution of KCl (75 mM) at 37°C for 15 min and fixed in three changes of methanol:acetic acid (3:1). The cell suspension is dropped onto a glass slide and air dried. The extended cDNA (or genomic DNA obtainable therefrom) is labeled with biotin-16  
5 dUTP by nick translation according to the manufacturer's instructions (Bethesda Research Laboratories, Bethesda, MD), purified using a Sephadex G-50 column (Pharmacia, Upssala, Sweden) and precipitated. Just prior to hybridization, the DNA pellet is dissolved in hybridization buffer (50% formamide, 2 X SSC, 10% dextran sulfate, 1 mg/ml sonicated salmon sperm DNA, pH 7) and the probe is denatured at 70°C for 5-10  
10 min.

Slides kept at -20°C are treated for 1 h at 37°C with RNase A (100 µg/ml), rinsed three times in 2 X SSC and dehydrated in an ethanol series. Chromosome preparations are denatured in 70% formamide, 2 X SSC for 2 min at 70°C, then dehydrated at 4°C. The slides are treated with proteinase K (10 µg/100 ml in 20 mM Tris-HCl, 2 mM CaCl<sub>2</sub>) at  
15 37°C for 8 min and dehydrated. The hybridization mixture containing the probe is placed on the slide, covered with a coverslip, sealed with rubber cement and incubated overnight in a humid chamber at 37°C. After hybridization and post-hybridization washes, the biotinylated probe is detected by avidin-FITC and amplified with additional layers of biotinylated goat anti-avidin and avidin-FITC. For chromosomal localization, fluorescent  
20 R-bands are obtained as previously described (Cherif et al., *supra*). The slides are observed under a LEICA fluorescence microscope (DMRXA). Chromosomes are counterstained with propidium iodide and the fluorescent signal of the probe appears as two symmetrical yellow-green spots on both chromatids of the fluorescent R-band chromosome (red). Thus, a particular extended cDNA (or genomic DNA obtainable  
25 therefrom) may be localized to a particular cytogenetic R-band on a given chromosome.

Once the extended cDNAs (or genomic DNAs obtainable therefrom) have been assigned to particular chromosomes using the techniques described in Examples 49-51 above, they may be utilized to construct a high resolution map of the chromosomes on which they are located or to identify the chromosomes in a sample.



## EXAMPLE 52

### Use of Extended cDNAs to Construct or Expand Chromosome Maps

Chromosome mapping involves assigning a given unique sequence to a particular chromosome as described above. Once the unique sequence has been mapped to a given chromosome, it is ordered relative to other unique sequences located on the same chromosome. One approach to chromosome mapping utilizes a series of yeast artificial chromosomes (YACs) bearing several thousand long inserts derived from the chromosomes of the organism from which the extended cDNAs (or genomic DNAs obtainable therefrom) are obtained. This approach is described in Ramaiah Nagaraja et al. **Genome Research** 7:210-222, March 1997. Briefly, in this approach each chromosome is broken into overlapping pieces which are inserted into the YAC vector. The YAC inserts are screened using PCR or other methods to determine whether they include the extended cDNA (or genomic DNA obtainable therefrom) whose position is to be determined. Once an insert has been found which includes the extended cDNA (or genomic DNA obtainable therefrom), the insert can be analyzed by PCR or other methods to determine whether the insert also contains other sequences known to be on the chromosome or in the region from which the extended cDNA (or genomic DNA obtainable therefrom) was derived. This process can be repeated for each insert in the YAC library to determine the location of each of the extended cDNAs (or genomic DNAs obtainable therefrom) relative to one another and to other known chromosomal markers. In this way, a high resolution map of the distribution of numerous unique markers along each of the organisms chromosomes may be obtained.

As described in Example 53 below extended cDNAs (or genomic DNAs obtainable therefrom) may also be used to identify genes associated with a particular phenotype, such as hereditary disease or drug response.

## EXAMPLE 53

### Identification of genes associated with hereditary diseases or drug response

This example illustrates an approach useful for the association of extended cDNAs (or genomic DNAs obtainable therefrom) with particular phenotypic characteristics. In this example, a particular extended cDNA (or genomic DNA obtainable therefrom) is used as a test probe to associate that extended cDNA (or genomic DNA obtainable therefrom) with a particular phenotypic characteristic.

Extended cDNAs (or genomic DNAs obtainable therefrom) are mapped to a particular location on a human chromosome using techniques such as those described in Examples 49 and 50 or other techniques known in the art. A search of Mendelian Inheritance in Man (V. McKusick, **Mendelian Inheritance in Man** (available on line through Johns Hopkins University Welch Medical Library) reveals the region of the human chromosome which contains the extended cDNA (or genomic DNA obtainable therefrom) to be a very gene rich region containing several known genes and several diseases or phenotypes for which genes have not been identified. The gene corresponding to this extended cDNA (or genomic DNA obtainable therefrom) thus becomes an immediate candidate for each of these genetic diseases.

Cells from patients with these diseases or phenotypes are isolated and expanded in culture. PCR primers from the extended cDNA (or genomic DNA obtainable therefrom) are used to screen genomic DNA, mRNA or cDNA obtained from the patients. Extended cDNAs (or genomic DNAs obtainable therefrom) that are not amplified in the patients can be positively associated with a particular disease by further analysis. Alternatively, the PCR analysis may yield fragments of different lengths when the samples are derived from an individual having the phenotype associated with the disease than when the sample is derived from a healthy individual, indicating that the gene containing the extended cDNA may be responsible for the genetic disease.

#### **VI. Use of Extended cDNAs (or genomic DNAs obtainable therefrom) to Construct Vectors**

The present extended cDNAs (or genomic DNAs obtainable therefrom) may also be used to construct secretion vectors capable of directing the secretion of the proteins encoded by genes inserted in the vectors. Such secretion vectors may facilitate the

purification or enrichment of the proteins encoded by genes inserted therein by reducing the number of background proteins from which the desired protein must be purified or enriched. Exemplary secretion vectors are described in Example 54 below.

#### EXAMPLE 54

##### 5                                    Construction of Secretion Vectors

The secretion vectors of the present invention include a promoter capable of directing gene expression in the host cell, tissue, or organism of interest. Such promoters include the Rous Sarcoma Virus promoter, the SV40 promoter, the human cytomegalovirus promoter, and other promoters familiar to those skilled in the art.

10                    A signal sequence from an extended cDNA (or genomic DNA obtainable therefrom), such as one of the signal sequences in SEQ ID NOs: 40-140 and 242-377 as defined in Table IV above, is operably linked to the promoter such that the mRNA transcribed from the promoter will direct the translation of the signal peptide. The host cell, tissue, or organism may be any cell, tissue, or organism which recognizes the signal  
15                    peptide encoded by the signal sequence in the extended cDNA (or genomic DNA obtainable therefrom). Suitable hosts include mammalian cells, tissues or organisms, avian cells, tissues, or organisms, insect cells, tissues or organisms, or yeast.

In addition, the secretion vector contains cloning sites for inserting genes encoding the proteins which are to be secreted. The cloning sites facilitate the cloning of the insert  
20                    gene in frame with the signal sequence such that a fusion protein in which the signal peptide is fused to the protein encoded by the inserted gene is expressed from the mRNA transcribed from the promoter. The signal peptide directs the extracellular secretion of the fusion protein.

The secretion vector may be DNA or RNA and may integrate into the  
25                    chromosome of the host, be stably maintained as an extrachromosomal replicon in the host, be an artificial chromosome, or be transiently present in the host. Many nucleic acid backbones suitable for use as secretion vectors are known to those skilled in the art, including retroviral vectors, SV40 vectors, Bovine Papilloma Virus vectors, yeast integrating plasmids, yeast episomal plasmids, yeast artificial chromosomes, human

artificial chromosomes, P element vectors, baculovirus vectors, or bacterial plasmids capable of being transiently introduced into the host.

The secretion vector may also contain a polyA signal such that the polyA signal is located downstream of the gene inserted into the secretion vector.

5        After the gene encoding the protein for which secretion is desired is inserted into the secretion vector, the secretion vector is introduced into the host cell, tissue, or organism using calcium phosphate precipitation, DEAE-Dextran, electroporation, liposome-mediated transfection, viral particles or as naked DNA. The protein encoded by the inserted gene is then purified or enriched from the supernatant using conventional  
10 techniques such as ammonium sulfate precipitation, immunoprecipitation, immunochromatography, size exclusion chromatography, ion exchange chromatography, and hplc. Alternatively, the secreted protein may be in a sufficiently enriched or pure state in the supernatant or growth media of the host to permit it to be used for its intended purpose without further enrichment.

15        The signal sequences may also be inserted into vectors designed for gene therapy. In such vectors, the signal sequence is operably linked to a promoter such that mRNA transcribed from the promoter encodes the signal peptide. A cloning site is located downstream of the signal sequence such that a gene encoding a protein whose secretion is desired may readily be inserted into the vector and fused to the signal sequence. The  
20 vector is introduced into an appropriate host cell. The protein expressed from the promoter is secreted extracellularly, thereby producing a therapeutic effect.

25        The extended cDNAs or 5' ESTs may also be used to clone sequences located upstream of the extended cDNAs or 5' ESTs which are capable of regulating gene expression, including promoter sequences, enhancer sequences, and other upstream sequences which influence transcription or translation levels. Once identified and cloned, these upstream regulatory sequences may be used in expression vectors designed to direct the expression of an inserted gene in a desired spatial, temporal, developmental, or quantitative fashion. Example 55 describes a method for cloning sequences upstream of the extended cDNAs or 5' ESTs.

## EXAMPLE 55

### Use of Extended cDNAs or 5' ESTs to Clone Upstream

#### Sequences from Genomic DNA

Sequences derived from extended cDNAs or 5' ESTs may be used to isolate the  
5 promoters of the corresponding genes using chromosome walking techniques. In one  
chromosome walking technique, which utilizes the GenomeWalker™ kit available from  
Clontech, five complete genomic DNA samples are each digested with a different  
restriction enzyme which has a 6 base recognition site and leaves a blunt end. Following  
digestion, oligonucleotide adapters are ligated to each end of the resulting genomic DNA  
10 fragments.

For each of the five genomic DNA libraries, a first PCR reaction is performed  
according to the manufacturer's instructions (which are incorporated herein by reference)  
using an outer adaptor primer provided in the kit and an outer gene specific primer. The  
gene specific primer should be selected to be specific for the extended cDNA or 5' EST of  
15 interest and should have a melting temperature, length, and location in the extended  
cDNA or ' EST which is consistent with its use in PCR reactions. Each first PCR reaction  
contains 5ng of genomic DNA, 5 µl of 10X Tth reaction buffer, 0.2 mM of each dNTP,  
0.2 µM each of outer adaptor primer and outer gene specific primer, 1.1 mM of  
Mg(OAc)<sub>2</sub>, and 1 µl of the Tth polymerase 50X mix in a total volume of 50 µl. The  
20 reaction cycle for the first PCR reaction is as follows: 1 min @ 94°C / 2 sec @ 94°C, 3  
min @ 72°C (7 cycles) / 2 sec @ 94°C, 3 min @ 67°C (32 cycles) / 5 min @ 67°C.

The product of the first PCR reaction is diluted and used as a template for a second  
PCR reaction according to the manufacturer's instructions using a pair of nested primers  
which are located internally on the amplicon resulting from the first PCR reaction. For  
25 example, 5 µl of the reaction product of the first PCR reaction mixture may be diluted 180  
times. Reactions are made in a 50 µl volume having a composition identical to that of the  
first PCR reaction except the nested primers are used. The first nested primer is specific  
for the adaptor, and is provided with the GenomeWalker™ kit. The second nested primer  
is specific for the particular extended cDNA or 5' EST for which the promoter is to be

cloned and should have a melting temperature, length, and location in the extended cDNA or 5' EST which is consistent with its use in PCR reactions. The reaction parameters of the second PCR reaction are as follows: 1 min @ 94°C / 2 sec @ 94°C, 3 min @ 72°C (6 cycles) / 2 sec @ 94°C, 3 min @ 67°C (25 cycles) / 5 min @ 67°C

5           The product of the second PCR reaction is purified, cloned, and sequenced using standard techniques. Alternatively, two or more human genomic DNA libraries can be constructed by using two or more restriction enzymes. The digested genomic DNA is cloned into vectors which can be converted into single stranded, circular, or linear DNA. A biotinylated oligonucleotide comprising at least 15 nucleotides from the extended  
10 cDNA or 5' EST sequence is hybridized to the single stranded DNA. Hybrids between the biotinylated oligonucleotide and the single stranded DNA containing the extended cDNA or EST sequence are isolated as described in Example 29 above. Thereafter, the single stranded DNA containing the extended cDNA or EST sequence is released from the beads and converted into double stranded DNA using a primer specific for the extended cDNA  
15 or 5' EST sequence or a primer corresponding to a sequence included in the cloning vector. The resulting double stranded DNA is transformed into bacteria. DNAs containing the 5' EST or extended cDNA sequences are identified by colony PCR or colony hybridization.

20           Once the upstream genomic sequences have been cloned and sequenced as described above, prospective promoters and transcription start sites within the upstream sequences may be identified by comparing the sequences upstream of the extended cDNAs or 5' ESTs with databases containing known transcription start sites, transcription factor binding sites, or promoter sequences.

25           In addition, promoters in the upstream sequences may be identified using promoter reporter vectors as described in Example 56.

## **EXAMPLE 56**

### **Identification of Promoters in Cloned Upstream Sequences**

The genomic sequences upstream of the extended cDNAs or 5' ESTs are cloned into a suitable promoter reporter vector, such as the pSEAP-Basic, pSEAP-Enhancer,

p $\beta$ gal-Basic, p $\beta$ gal-Enhancer, or pEGFP-1 Promoter Reporter vectors available from Clontech. Briefly, each of these promoter reporter vectors include multiple cloning sites positioned upstream of a reporter gene encoding a readily assayable protein such as secreted alkaline phosphatase,  $\beta$  galactosidase, or green fluorescent protein. The sequences upstream of the extended cDNAs or 5' ESTs are inserted into the cloning sites upstream of the reporter gene in both orientations and introduced into an appropriate host cell. The level of reporter protein is assayed and compared to the level obtained from a vector which lacks an insert in the cloning site. The presence of an elevated expression level in the vector containing the insert with respect to the control vector indicates the presence of a promoter in the insert. If necessary, the upstream sequences can be cloned into vectors which contain an enhancer for augmenting transcription levels from weak promoter sequences. A significant level of expression above that observed with the vector lacking an insert indicates that a promoter sequence is present in the inserted upstream sequence.

Appropriate host cells for the promoter reporter vectors may be chosen based on the results of the above described determination of expression patterns of the extended cDNAs and ESTs. For example, if the expression pattern analysis indicates that the mRNA corresponding to a particular extended cDNA or 5' EST is expressed in fibroblasts, the promoter reporter vector may be introduced into a human fibroblast cell line.

Promoter sequences within the upstream genomic DNA may be further defined by constructing nested deletions in the upstream DNA using conventional techniques such as Exonuclease III digestion. The resulting deletion fragments can be inserted into the promoter reporter vector to determine whether the deletion has reduced or obliterated promoter activity. In this way, the boundaries of the promoters may be defined. If desired, potential individual regulatory sites within the promoter may be identified using site directed mutagenesis or linker scanning to obliterate potential transcription factor binding sites within the promoter individually or in combination. The effects of these

mutations on transcription levels may be determined by inserting the mutations into the cloning sites in the promoter reporter vectors.

## EXAMPLE 57

### Cloning and Identification of Promoters

5           Using the method described in Example 55 above with 5' ESTs, sequences upstream of several genes were obtained. Using the primer pairs GGG AAG ATG GAG ATA GTA TTG CCT G (SEQ ID NO:29) and CTG CCA TGT ACA TGA TAG AGA GAT TC (SEQ ID NO:30), the promoter having the internal designation P13H2 (SEQ ID NO:31) was obtained.

10           Using the primer pairs GTA CCA GGGG ACT GTG ACC ATT GC (SEQ ID NO:32) and CTG TGA CCA TTG CTC CCA AGA GAG (SEQ ID NO:33), the promoter having the internal designation P15B4 (SEQ ID NO:34) was obtained.

          Using the primer pairs CTG GGA TGG AAG GCA CGG TA (SEQ ID NO:35) and GAG ACC ACA CAG CTA GAC AA (SEQ ID NO:36), the promoter having the  
15           internal designation P29B6 (SEQ ID NO:37) was obtained.

          Figure 8 provides a schematic description of the promoters isolated and the way they are assembled with the corresponding 5' tags. The upstream sequences were screened for the presence of motifs resembling transcription factor binding sites or known transcription start sites using the computer program MatInspector release 2.0, August  
20           1996.

          Figure 9 describes the transcription factor binding sites present in each of these promoters. The columns labeled matrice provides the name of the MatInspector matrix used. The column labeled position provides the 5' position of the promoter site. Numeration of the sequence starts from the transcription site as determined by matching  
25           the genomic sequence with the 5' EST sequence. The column labeled "orientation" indicates the DNA strand on which the site is found, with the + strand being the coding strand as determined by matching the genomic sequence with the sequence of the 5' EST. The column labeled "score" provides the MatInspector score found for this site. The



column labeled "length" provides the length of the site in nucleotides. The column labeled "sequence" provides the sequence of the site found.

The promoters and other regulatory sequences located upstream of the extended cDNAs or 5' ESTs may be used to design expression vectors capable of directing the expression of an inserted gene in a desired spatial, temporal, developmental, or quantitative manner. A promoter capable of directing the desired spatial, temporal, developmental, and quantitative patterns may be selected using the results of the expression analysis described in Example 26 above. For example, if a promoter which confers a high level of expression in muscle is desired, the promoter sequence upstream of an extended cDNA or 5' EST derived from an mRNA which is expressed at a high level in muscle, as determined by the method of Example 26, may be used in the expression vector.

Preferably, the desired promoter is placed near multiple restriction sites to facilitate the cloning of the desired insert downstream of the promoter, such that the promoter is able to drive expression of the inserted gene. The promoter may be inserted in conventional nucleic acid backbones designed for extrachromosomal replication, integration into the host chromosomes or transient expression. Suitable backbones for the present expression vectors include retroviral backbones, backbones from eukaryotic episomes such as SV40 or Bovine Papilloma Virus, backbones from bacterial episomes, or artificial chromosomes.

Preferably, the expression vectors also include a polyA signal downstream of the multiple restriction sites for directing the polyadenylation of mRNA transcribed from the gene inserted into the expression vector.

Following the identification of promoter sequences using the procedures of Examples 55-57, proteins which interact with the promoter may be identified as described in Example 58 below.

### **EXAMPLE 58**

#### **Identification of Proteins Which Interact with Promoter Sequences, Upstream Regulatory Sequences, or mRNA**

Sequences within the promoter region which are likely to bind transcription factors may be identified by homology to known transcription factor binding sites or through conventional mutagenesis or deletion analyses of reporter plasmids containing the promoter sequence. For example, deletions may be made in a reporter plasmid containing the promoter sequence of interest operably linked to an assayable reporter gene. The reporter plasmids carrying various deletions within the promoter region are transfected into an appropriate host cell and the effects of the deletions on expression levels is assessed. Transcription factor binding sites within the regions in which deletions reduce expression levels may be further localized using site directed mutagenesis, linker scanning analysis, or other techniques familiar to those skilled in the art. Nucleic acids encoding proteins which interact with sequences in the promoter may be identified using one-hybrid systems such as those described in the manual accompanying the Matchmaker One-Hybrid System kit available from Clontech (Catalog No. K1603-1), the disclosure of which is incorporated herein by reference. Briefly, the Matchmaker One-hybrid system is used as follows. The target sequence for which it is desired to identify binding proteins is cloned upstream of a selectable reporter gene and integrated into the yeast genome. Preferably, multiple copies of the target sequences are inserted into the reporter plasmid in tandem.

A library comprised of fusions between cDNAs to be evaluated for the ability to bind to the promoter and the activation domain of a yeast transcription factor, such as GAL4, is transformed into the yeast strain containing the integrated reporter sequence. The yeast are plated on selective media to select cells expressing the selectable marker linked to the promoter sequence. The colonies which grow on the selective media contain genes encoding proteins which bind the target sequence. The inserts in the genes encoding the fusion proteins are further characterized by sequencing. In addition, the inserts may be inserted into expression vectors or in vitro transcription vectors. Binding of the polypeptides encoded by the inserts to the promoter DNA may be confirmed by techniques familiar to those skilled in the art, such as gel shift analysis or DNase protection analysis.

## VII. Use of Extended cDNAs (or Genomic DNAs Obtainable Therefrom) in Gene Therapy

The present invention also comprises the use of extended cDNAs (or genomic DNAs obtainable therefrom) in gene therapy strategies, including antisense and triple helix strategies as described in Examples 57 and 58 below. In antisense approaches, nucleic acid sequences complementary to an mRNA are hybridized to the mRNA intracellularly, thereby blocking the expression of the protein encoded by the mRNA. The antisense sequences may prevent gene expression through a variety of mechanisms. For example, the antisense sequences may inhibit the ability of ribosomes to translate the mRNA. Alternatively, the antisense sequences may block transport of the mRNA from the nucleus to the cytoplasm, thereby limiting the amount of mRNA available for translation. Another mechanism through which antisense sequences may inhibit gene expression is by interfering with mRNA splicing. In yet another strategy, the antisense nucleic acid may be incorporated in a ribozyme capable of specifically cleaving the target mRNA.

### EXAMPLE 59

#### Preparation and Use of Antisense Oligonucleotides

The antisense nucleic acid molecules to be used in gene therapy may be either DNA or RNA sequences. They may comprise a sequence complementary to the sequence of the extended cDNA (or genomic DNA obtainable therefrom). The antisense nucleic acids should have a length and melting temperature sufficient to permit formation of an intracellular duplex having sufficient stability to inhibit the expression of the mRNA in the duplex. Strategies for designing antisense nucleic acids suitable for use in gene therapy are disclosed in Green et al., **Ann. Rev. Biochem.** 55:569-597 (1986) and Izant and Weintraub, **Cell** 36:1007-1015 (1984), which are hereby incorporated by reference.

In some strategies, antisense molecules are obtained from a nucleotide sequence encoding a protein by reversing the orientation of the coding region with respect to a promoter so as to transcribe the opposite strand from that which is normally transcribed in the cell. The antisense molecules may be transcribed using in vitro transcription systems

such as those which employ T7 or SP6 polymerase to generate the transcript. Another approach involves transcription of the antisense nucleic acids in vivo by operably linking DNA containing the antisense sequence to a promoter in an expression vector.

Alternatively, oligonucleotides which are complementary to the strand normally transcribed in the cell may be synthesized in vitro. Thus, the antisense nucleic acids are complementary to the corresponding mRNA and are capable of hybridizing to the mRNA to create a duplex. In some embodiments, the antisense sequences may contain modified sugar phosphate backbones to increase stability and make them less sensitive to RNase activity. Examples of modifications suitable for use in antisense strategies are described by Rossi et al., **Pharmacol. Ther.** 50(2):245-254, (1991).

Various types of antisense oligonucleotides complementary to the sequence of the extended cDNA (or genomic DNA obtainable therefrom) may be used. In one preferred embodiment, stable and semi-stable antisense oligonucleotides described in International Application No. PCT WO94/23026, hereby incorporated by reference, are used. In these molecules, the 3' end or both the 3' and 5' ends are engaged in intramolecular hydrogen bonding between complementary base pairs. These molecules are better able to withstand exonuclease attacks and exhibit increased stability compared to conventional antisense oligonucleotides.

In another preferred embodiment, the antisense oligodeoxynucleotides against herpes simplex virus types 1 and 2 described in International Application No. WO 95/04141, hereby incorporated by reference, are used.

In yet another preferred embodiment, the covalently cross-linked antisense oligonucleotides described in International Application No. WO 96/31523, hereby incorporated by reference, are used. These double- or single-stranded oligonucleotides comprise one or more, respectively, inter- or intra-oligonucleotide covalent cross-linkages, wherein the linkage consists of an amide bond between a primary amine group of one strand and a carboxyl group of the other strand or of the same strand, respectively, the primary amine group being directly substituted in the 2' position of the strand nucleotide monosaccharide ring, and the carboxyl group being carried by an aliphatic spacer group

substituted on a nucleotide or nucleotide analog of the other strand or the same strand, respectively.

The antisense oligodeoxynucleotides and oligonucleotides disclosed in International Application No. WO 92/18522, incorporated by reference, may also be used.

5 These molecules are stable to degradation and contain at least one transcription control recognition sequence which binds to control proteins and are effective as decoys therefor. These molecules may contain "hairpin" structures, "dumbbell" structures, "modified dumbbell" structures, "cross-linked" decoy structures and "loop" structures.

10 In another preferred embodiment, the cyclic double-stranded oligonucleotides described in European Patent Application No. 0 572 287 A2, hereby incorporated by reference are used. These ligated oligonucleotide "dumbbells" contain the binding site for a transcription factor and inhibit expression of the gene under control of the transcription factor by sequestering the factor.

15 Use of the closed antisense oligonucleotides disclosed in International Application No. WO 92/19732, hereby incorporated by reference, is also contemplated. Because these molecules have no free ends, they are more resistant to degradation by exonucleases than are conventional oligonucleotides. These oligonucleotides may be multifunctional, interacting with several regions which are not adjacent to the target mRNA.

20 The appropriate level of antisense nucleic acids required to inhibit gene expression may be determined using in vitro expression analysis. The antisense molecule may be introduced into the cells by diffusion, injection, infection or transfection using procedures known in the art. For example, the antisense nucleic acids can be introduced into the body as a bare or naked oligonucleotide, oligonucleotide encapsulated in lipid, oligonucleotide sequence encapsidated by viral protein, or as an oligonucleotide operably linked to a  
25 promoter contained in an expression vector. The expression vector may be any of a variety of expression vectors known in the art, including retroviral or viral vectors, vectors capable of extrachromosomal replication, or integrating vectors. The vectors may be DNA or RNA.

The antisense molecules are introduced onto cell samples at a number of different concentrations preferably between  $1 \times 10^{-10} \text{M}$  to  $1 \times 10^{-4} \text{M}$ . Once the minimum concentration that can adequately control gene expression is identified, the optimized dose is translated into a dosage suitable for use in vivo. For example, an inhibiting concentration in culture of  $1 \times 10^{-7}$  translates into a dose of approximately 0.6 mg/kg bodyweight. Levels of oligonucleotide approaching 100 mg/kg bodyweight or higher may be possible after testing the toxicity of the oligonucleotide in laboratory animals. It is additionally contemplated that cells from the vertebrate are removed, treated with the antisense oligonucleotide, and reintroduced into the vertebrate.

It is further contemplated that the antisense oligonucleotide sequence is incorporated into a ribozyme sequence to enable the antisense to specifically bind and cleave its target mRNA. For technical applications of ribozyme and antisense oligonucleotides see Rossi et al., *supra*.

In a preferred application of this invention, the polypeptide encoded by the gene is first identified, so that the effectiveness of antisense inhibition on translation can be monitored using techniques that include but are not limited to antibody-mediated tests such as RIAs and ELISA, functional assays, or radiolabeling.

The extended cDNAs of the present invention (or genomic DNAs obtainable therefrom) may also be used in gene therapy approaches based on intracellular triple helix formation. Triple helix oligonucleotides are used to inhibit transcription from a genome. They are particularly useful for studying alterations in cell activity as it is associated with a particular gene. The extended cDNAs (or genomic DNAs obtainable therefrom) of the present invention or, more preferably, a portion of those sequences, can be used to inhibit gene expression in individuals having diseases associated with expression of a particular gene. Similarly, a portion of the extended cDNA (or genomic DNA obtainable therefrom) can be used to study the effect of inhibiting transcription of a particular gene within a cell. Traditionally, homopurine sequences were considered the most useful for triple helix strategies. However, homopyrimidine sequences can also inhibit gene expression. Such homopyrimidine oligonucleotides bind to the major groove at

homopurine:homopyrimidine sequences. Thus, both types of sequences from the extended cDNA or from the gene corresponding to the extended cDNA are contemplated within the scope of this invention.

#### EXAMPLE 60

##### 5                    Preparation and use of Triple Helix Probes

          The sequences of the extended cDNAs (or genomic DNAs obtainable therefrom) are scanned to identify 10-mer to 20-mer homopyrimidine or homopurine stretches which could be used in triple-helix based strategies for inhibiting gene expression. Following identification of candidate homopyrimidine or homopurine stretches, their efficiency in  
10    inhibiting gene expression is assessed by introducing varying amounts of oligonucleotides containing the candidate sequences into tissue culture cells which normally express the target gene. The oligonucleotides may be prepared on an oligonucleotide synthesizer or they may be purchased commercially from a company specializing in custom oligonucleotide synthesis, such as GENSET, Paris, France.

15            The oligonucleotides may be introduced into the cells using a variety of methods known to those skilled in the art, including but not limited to calcium phosphate precipitation, DEAE-Dextran, electroporation, liposome-mediated transfection or native uptake.

          Treated cells are monitored for altered cell function or reduced gene expression  
20    using techniques such as Northern blotting, RNase protection assays, or PCR based strategies to monitor the transcription levels of the target gene in cells which have been treated with the oligonucleotide. The cell functions to be monitored are predicted based upon the homologies of the target gene corresponding to the extended cDNA from which the oligonucleotide was derived with known gene sequences that have been associated  
25    with a particular function. The cell functions can also be predicted based on the presence of abnormal physiologies within cells derived from individuals with a particular inherited disease, particularly when the extended cDNA is associated with the disease using techniques described in Example 53.

The oligonucleotides which are effective in inhibiting gene expression in tissue culture cells may then be introduced in vivo using the techniques described above and in Example 59 at a dosage calculated based on the in vitro results, as described in Example 59.

5 In some embodiments, the natural (beta) anomers of the oligonucleotide units can be replaced with alpha anomers to render the oligonucleotide more resistant to nucleases. Further, an intercalating agent such as ethidium bromide, or the like, can be attached to the 3' end of the alpha oligonucleotide to stabilize the triple helix. For information on the generation of oligonucleotides suitable for triple helix formation see Griffin et al. (**Science**  
10 **245**:967-971 (1989), which is hereby incorporated by this reference).

### EXAMPLE 61

#### Use of Extended cDNAs to Express an Encoded Protein in a Host Organism

The extended cDNAs of the present invention may also be used to express an encoded protein in a host organism to produce a beneficial effect. In such procedures, the  
15 encoded protein may be transiently expressed in the host organism or stably expressed in the host organism. The encoded protein may have any of the activities described above. The encoded protein may be a protein which the host organism lacks or, alternatively, the encoded protein may augment the existing levels of the protein in the host organism.

A full length extended cDNA encoding the signal peptide and the mature protein,  
20 or an extended cDNA encoding only the mature protein is introduced into the host organism. The extended cDNA may be introduced into the host organism using a variety of techniques known to those of skill in the art. For example, the extended cDNA may be injected into the host organism as naked DNA such that the encoded protein is expressed in the host organism, thereby producing a beneficial effect.

25 Alternatively, the extended cDNA may be cloned into an expression vector downstream of a promoter which is active in the host organism. The expression vector may be any of the expression vectors designed for use in gene therapy, including viral or retroviral vectors.



The expression vector may be directly introduced into the host organism such that the encoded protein is expressed in the host organism to produce a beneficial effect. In another approach, the expression vector may be introduced into cells *in vitro*. Cells containing the expression vector are thereafter selected and introduced into the host organism, where they express the encoded protein to produce a beneficial effect.

## EXAMPLE 62

### Use Of Signal Peptides Encoded By 5' Ests Or Sequences Obtained Therefrom To Import Proteins Into Cells

The short core hydrophobic region (h) of signal peptides encoded by the 5'ESTS or extended cDNAs derived from the 5'ESTs of the present invention may also be used as a carrier to import a peptide or a protein of interest, so-called cargo, into tissue culture cells (Lin *et al.*, *J. Biol. Chem.*, **270**: 14225-14258 (1995); Du *et al.*, *J. Peptide Res.*, **51**: 235-243 (1998); Rojas *et al.*, *Nature Biotech.*, **16**: 370-375 (1998)).

When cell permeable peptides of limited size (approximately up to 25 amino acids) are to be translocated across cell membrane, chemical synthesis may be used in order to add the h region to either the C-terminus or the N-terminus to the cargo peptide of interest. Alternatively, when longer peptides or proteins are to be imported into cells, nucleic acids can be genetically engineered, using techniques familiar to those skilled in the art, in order to link the extended cDNA sequence encoding the h region to the 5' or the 3' end of a DNA sequence coding for a cargo polypeptide. Such genetically engineered nucleic acids are then translated either *in vitro* or *in vivo* after transfection into appropriate cells, using conventional techniques to produce the resulting cell permeable polypeptide. Suitable hosts cells are then simply incubated with the cell permeable polypeptide which is then translocated across the membrane.

This method may be applied to study diverse intracellular functions and cellular processes. For instance, it has been used to probe functionally relevant domains of intracellular proteins and to examine protein-protein interactions involved in signal transduction pathways (Lin *et al.*, *supra*; Lin *et al.*, *J. Biol. Chem.*, **271**: 5305-5308 (1996); Rojas *et al.*, *J. Biol. Chem.*, **271**: 27456-27461 (1996); Liu *et al.*, *Proc. Natl. Acad.*

*Sci. USA*, **93**: 11819-11824 (1996); Rojas *et al.*, *Bioch. Biophys. Res. Commun.*, **234**: 675-680 (1997)).

Such techniques may be used in cellular therapy to import proteins producing therapeutic effects. For instance, cells isolated from a patient may be treated with imported therapeutic proteins and then re-introduced into the host organism.

Alternatively, the h region of signal peptides of the present invention could be used in combination with a nuclear localization signal to deliver nucleic acids into cell nucleus. Such oligonucleotides may be antisense oligonucleotides or oligonucleotides designed to form triple helixes, as described in examples 59 and 60 respectively, in order to inhibit processing and maturation of a target cellular RNA.

### EXAMPLE 63

#### Reassembling & Resequencing of Clones

Full length cDNA clones obtained by the procedure described in Example 27 were double-sequenced. These sequences were assembled and the resulting consensus sequences were then reanalyzed. Open reading frames were reassigned following essentially the same process as the one described in Example 27.

After this reanalysis process a few abnormalities were revealed. The sequences presented in SEQ ID NOs: 47, 73, 79, 89, 91, 96, 126, 128, 134, and 139 are apparently unlikely to be genuine full length cDNAs. These clones are missing a stop codon and are thus more probably 3' truncated cDNA sequences. Similarly, the sequences presented in SEQ ID NOs: 45, 50, 54, 57, 73, 74, 89, 92, 95, 98, 126, 129, 130, 131 and 139 may also not be genuine full length cDNAs based on homology studies with existing protein sequences. Although both of these sequences encode a potential start methionine each could represent a 5' truncated cDNA.

In addition, SEQ ID NO: 115 was found to be an alternatively spliced transcript and the identities of some of the bases in SEQ ID NO: 131 were corrected.

Finally, after the reassignment of open reading frames for the clones, new open reading frames were chosen in some instances. For example, in the case of SEQ ID NOs: 41, 47, 50, 52, 54-56, 58, 59, 61, 74, 75, 79, 84, 89, 91, 92, 96, 98, 103, 105, 106,

126, 129, 131, and 133 the new open reading frames were no longer predicted to contain a signal peptide.

As discussed above, Table IV provides the sequence identification numbers of the extended cDNAs of the present invention, the locations of the full coding sequences in SEQ ID NOs: 40-140 and 242-377 (i.e. the nucleotides encoding both the signal peptide and the mature protein, listed under the heading FCS location in Table IV), the locations of the nucleotides in SEQ ID NOs: 40-140 and 242-377 which encode the signal peptides (listed under the heading SigPep Location in Table IV), the locations of the nucleotides in SEQ ID NOs: 40-140 and 242-377 which encode the mature proteins generated by cleavage of the signal peptides (listed under the heading Mature Polypeptide Location in Table IV), the locations in SEQ ID NOs: 40-140 and 242-377 of stop codons (listed under the heading Stop Codon Location in Table IV) the locations in SEQ ID NOs: 40-140 and 242-377 of polyA signals (listed under the heading g PolyA Signal Location in Table IV) and the locations of polyA sites (listed under the heading PolyA Site Location in Table IV).

As discussed above, Table V lists the sequence identification numbers of the polypeptides of SEQ ID NOs: 141-241 and 378-513, the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the full length polypeptide (second column), the locations of the amino acid residues of SEQ ID NOs: 141-241 and 378-513 in the signal peptides (third column), and the locations of the amino acid residues of SEQ ID NOs: 141-241 and 379-513 in the mature polypeptide created by cleaving the signal peptide from the full length polypeptide (fourth column). In Table V, and in the appended sequence listing, the first amino acid of the mature protein resulting from cleavage of the signal peptide is designated as amino acid number 1 and the first amino acid of the signal peptide is designated with the appropriate negative number, in accordance with the regulations governing sequence listings.

#### EXAMPLE 64

##### Functional Analysis of Predicted Protein Sequences

Following double-sequencing, new contigs were assembled for each of the extended cDNAs of the present invention and each was compared to known sequences available at the time of filing. These sequences originate from the following databases: Genbank (release 108 and daily releases up to October, 15, 1998), Genseq (release 32)  
5 PIR (release 53) and SwissProt (release 35). The predicted proteins of the present invention matching known proteins were further classified into 3 categories depending on the level of homology.

The first category contains proteins of the present invention exhibiting more than 70% identical amino acid residues on the whole length of the matched protein.  
10 They are clearly close homologues which most probably have the same function or a very similar function as the matched protein.

The second category contains proteins of the present invention exhibiting more remote homologies (40 to 70% over the whole protein) indicating that the protein of the present invention may have functions similar to those of the homologous protein.

15 The third category contains proteins exhibiting homology (90 to 100%) to a domain of a known protein indicating that the matched protein and the protein of the invention may share similar features.

It should be noted that the numbering of amino acids in the protein sequences discussed in Figures 10 to 15, and Table VIII, the first methionine encountered is designated as amino acid number 1. In the appended sequence listing, the first amino  
20 acid of the mature protein resulting from cleavage of the signal peptide is designated as amino acid -number 1, and the first amino acid of the signal peptide is designated with the appropriate negative number, in accordance with the regulations governing sequence listings.

25 In addition all of the corrected amino acid sequences (SEQ ED NOs: 141-241 and 378-513) were scanned for the presence of known protein signatures and motifs. This search was performed against the Prosite 15.0 database, using the Proscan software from the GCG package- Functional signatures and their locations are indicated in Table VIII.

### **A) Proteins which are closely related to known proteins**

#### **Protein of SEQ ID NO: 217 (internal designation 48-46-4-A11-CL 1 4 p)**

5 The protein of SEQ ID NO: 217 encoded by the extended cDNA SEQ ID NO: 116 isolated from lymphocyte shows complete identity to a human protein TFAR19 that may play a role in apoptosis (Genbank accession number AF014955, SEQ ID NO: 516) as shown by the alignment in figure 10.

10 Taken together, these data suggest that the protein of SEQ ID NO: 217 may be involved in the control of development and homeostasis. Thus, this protein may be useful in diagnosis and/or treating several types of disorders including, but not limited to, cancer, autoimmune disorders, viral infections such as AIDS, neurodegenerative disorders, osteoporosis.

#### **Proteins of SEQ ID NOs: 174, 175 and 232 (internal designation 45-54-1-G9)**

15 The proteins of SEQ ID NOs: 174, 175 and 232 encoded by the extended cDNAs SEQ ID NOs: 73, 74 and 131 respectively and isolated from lymphocytes shows complete extensive homologies to a human secreted protein (Genbank accession number W36955, SEQ ID NO: 517). As shown by the alignments of figure 11, the amino acid residues are identical to those of the 110 amino acid long matched protein except for positions 51 and 108-110 of the matched protein for the protein of SEQ ID  
20 NOs: 174, for positions 48, 94 and 108-110 of the matched protein of SEQ ID NOs: 175 and for positions 94, and 108-110 of the matched protein for the protein of SEQ ID NOs: 232. Proteins of SEQ ID NOs: 174 and 232 may represent alternative forms issued from alternative use of polyadenylation signals.

25 Taken together, these data suggest that the proteins of SEQ ID NOs: 174, 175 and 232 may play a role in cell proliferation and/or differentiation, in immune responses and/or in haematopoiesis. Thus, this protein or part therein, may be useful in diagnosing and treating several disorders including, but not limited to, cancer, immunological, haematological and/or inflammatory disorders. It may also be useful in

modulating the immune and inflammatory responses to infectious agents and/or to suppress graft rejection .

Proteins of SEQ ID NO: 231 (internal designation 47-4-4-C6-CL2 2)

5           The protein of SEQ ID NO: 231 encoded by the extended cDNA SEQ ID NO: 130 shows extensive homology with the human E25 protein (Genbank accession number AF038953, SEQ ID NO: 515). As shown by the alignments in figure 12, the amino acid residues are identical except for position 159 in the 263 amino acid long matched sequence. The matched protein might be involved in the development and  
10 differentiation of haematopoietic stem/progenitor cells. In addition, it is the human homologue of a murine protein thought to be involved in chondro-osteogenic differentiation and belonging to a novel multigene family of integral membrane proteins (Deleersnijder *et al*, *J. Biol. Chem.*, **271** : 19475-19482 (1996)).

15           The protein of invention contains two short segments from positions 1 to 21 and from 100 to 120 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, **10** : 685-686 (1994)). The first transmembrane domains matches exactly those predicted for the murine E25 protein.

20           Taken together, these data suggest that the protein of SEQ ID NO: 231 may be involved in cellular proliferation and differentiation. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer and embryogenesis disorders.

Protein of SEQ ID NO: 196 (internal designation 76-13-3-A9-CL1 2)

25           The protein of SEQ ID NO: 196 encoded by the extended cDNA SEQ ID NO: 95 shows extensive homology with the human seventransmembrane protein (Genbank accession number Y11395, SEQ ID NO: 518) and its murine homologue (Genbank accession number Y11550). As shown by the alignments in figure 13, the amino acid residues are identical except for position 174 in the 399 amino acid long human matched sequence. The matched protein potentially associated to stomatin may act as a

G-protein coupled receptor and is likely to be important for the signal transduction in neurons and haematopoietic cells (Mayer *et al*, *Biochem. Biophys. Acta.*, **1395** : 301-308 (1998)).

Taken together, these data suggest that the protein of SEQ ID NOs: 196 may be involved in signal transduction. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases cardiovascular disorders, hypertension, renal injury and repair and septic shock.

Protein of SEQ ID NO: 158 (internal designation 33-35-4-G1-CL1 2)

The protein of SEQ ID NOs: 158 encoded by the extended cDNA SEQ ID NO: 57 shows homology with the murine subunit 7a of the COP9 complex (Genbank accession number AF071316, SEQ ID NO: 519). As shown by the alignments in figure 14, the amino acid residues are identical except for positions 90, 172 and 247 in the 275 amino acid long matched sequence. This complex is highly conserved between mammals and higher plants where it has been shown to act as a repressor of photomorphogenesis All the components of the mammalian COP9 complex contain structural features also present in components of the proteasome regulatory complex and the translation initiation complex eIF3 complex, suggesting that the mammalian COP9 complex is an important cellular regulator modulating multiple signaling pathways (Wei *et al*, *Curr. Biol.*, **8** : 919-922 (1998)).

Taken together, these data suggest that the protein of SEQ ID NO: 158 may be involved in cellular signaling, probably as a subunit of the human COP9 complex. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases, cardiovascular disorders, hypertension, renal injury and repair and septic shock.

Protein of SEQ ID NO: 226 (internal designation 19-10-1-C2-CL1 3)

The protein of SEQ ID NO: 226 encoded by the extended cDNA SEQ ID NO: 125 shows homology with the bovine subunit B14.5B of the NADH-ubiquinone oxidoreductase complex (Arizmendi *et al*, *FEBS Lett.*, **313** : 80-84 (1992) and Swissprot accession -number Q02827, SEQ ID NO: 514). As shown by the alignments in figure 5 15, the amino acid residues are identical except for positions 3-4, 6-12, 32-34, 47, 53-55, 67 and 69-74 in the 120 amino acid long matched sequence. This complex is the first of four complexes located in the inner mitochondrial membrane and composing the mitochondrial electron transport chain. Complex I is involved in the dehydrogenation of NADH and the transportation of electrons to coenzyme Q. It is composed of 7 10 subunits encoded by the mitochondrial genome and 34 subunits encoded by the nuclear genome. It is also thought to play a role in the regulation of apoptosis and necrosis. Mitochondriocytopathies due to complex I deficiency are frequently encountered and affect tissues with a high energy demand such as brain (mental retardation, convulsions, movement disorders), heart (cardiomyopathy, conduction disorders), kidney (Fanconi 15 syndrome), skeletal muscle (exercise intolerance, muscle weakness, hypotonia) and/or eye (ophthalmoplegia, ptosis, cataract and retinopathy). For a review on complex I see Smeitink *et al.*, *Hum. Mol. Gent.*, **7** : 1573-1579 (1998).

Taken together, these data suggest that the protein of SEQ ID NO: 226 may be part of the mitochondrial energy-generating system, probably as a subunit of the 20 NADH-ubiquinone oxidoreductase complex. Thus, this protein or part therein, may be useful in diagnosing and/or treating several disorders including, but not limited to, brain disorders (mental retardation, convulsions, movement disorders), heart disorders (cardiomyopathy, conduction disorders), kidney disorders (Fanconi syndrome), skeletal muscle disorders (exercise intolerance, muscle weakness, hypotonia) and/or eye 25 disorders ophthalmoplegia, ptosis, cataract and retinopathy).

## **B) Proteins which are remotely related to proteins with known functions**

Proteins of SEQ ID NOs: 149, 150 and 211 (internal designation 27-1-2-B3-CL0 x p)



The proteins of SEQ ID NOs: 149, 150 and 211 encoded by the extended cDNAs SEQ ID NOs: 48, 49 and 110 respectively and found in, skeletal muscle shows homologies with T1/ST2 ligand polypeptide of either human (Genbank accession number U41804 and Genseq accession number WO9639) or rodent species (Genbank accession number U41805 and Genseq accession number WO9640). These polypeptides are thought to be cytokines that bind to the ST2 receptor, a member of the immunoglobulin family homologous to the interleukin-1 receptor and present on some lymphoma cells. They are predicted to be cell-surface proteins containing a short transmembrane domain. (Gayle *et al*, *J. Biol. Chem.*, **271** : 5784-5789 (1996)). Proteins of SEQ ID NOs: 149, 150 and 211 may represent alternative forms issued from alternative use of polyadenylation signals.

The protein of invention contains two short transmembrane segments from positions 5 to 25 and from 195 to 215 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, **10** :685-686 (1994)). The second transmembrane domain matches exactly those of the matched cell-surface protein.

Taken together, these data suggest that the protein of SEQ ID NOs: 149, 150 and 211 may act as a cytokine, thus may play a role in the regulation of cell growth and differentiation and/or in the regulation of the immune response. Thus, this protein or part therein, may be useful in diagnosing and treating several disorders including, but not limited to, cancer, immunological, haematological and/or inflammatory disorders. It may also be useful in modulating the immune and inflammatory responses to infectious agents such as HIV and/or to suppress graft rejection.

#### Protein of SEQ ID NO: 177 (internal designation 51-11-3-D5-CL1 3)

The protein SEQ ID NO: 177 found in testis encoded by the extended cDNA SEQ ID NO: 76 shows homologies to serine protease inhibitor proteins belonging to the pancreatic trypsin inhibitor family (Kunitz) such as the extracellular proteinase inhibitor named chelonianin (Swissprot accession number P00993). The characteristic PROSITE

signature of this family is conserved in the protein of the invention (positions 69 to 87) except for a drastic change of the last cysteine residue into an arginine residue.

Taken together, these data suggest that the protein of SEQ ID NO: 177 may be a protease inhibitor, probably of the Kunitz family. Thus, this protein or part therein, may be useful in diagnosing and treating several disorders including but not limited to, cancer and neurodegenerative disorders such as Alzheimer's disease.

#### Protein of SEQ ID NO: 146 (internal designation 26-27-3-D7-CL0 1)

The protein SEQ ID NO: 146 encoded by the extended cDNA SEQ ID NO: 45 shows homology to human apolipoprotein L (Genbank accession number AF019225). The matched protein is a secreted high density lipoprotein associated with apoA-I-containing lipoproteins which play a key role in reverse cholesterol transport.

Taken together, these data suggest that the protein of SEQ ID NO: 146 may play a role in lipid metabolism. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, hyperlipidemia, hypercholesterolemia, atherosclerosis, cardiovascular disorders such as, coronary heart disease, and neurodegenerative disorders such as Alzheimer's disease or dementia.

#### Protein of SEQ ID NO: 163 (internal designation 33-49-1-H4-CL1 1)

The protein SEQ ED NO: 163 encoded by the extended cDNA SEQ ID NO: 62 shows homology to the yeast autophagocytosis protein AUT1 (SwissProt accession number P40344). The matched protein is required for starvation-induced non-specific bulk transport of cytoplasmic proteins to the vacuole.

Taken together, these data suggest that the protein of SEQ ID NO: 163 may play a role in protein transport. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, autoimmune disorders and immune disorders due to dysfunction of antigen presentation.

### **C) Proteins homologous to a domain of a protein with known function**

Protein of SEQ ID NO: 214 (internal designation 33-28-4-D1-CL0 1 p)

The protein of SEQ ID NO: 214 encoded by the extended cDNA SEQ ID NO: 113 and expressed in adult brain shows extensive homology to part of the murine SHYC protein (Genbank accession number AF072697) which is expressed in the developing and embryonic nervous system as well as along the olfactory pathway in adult brains (Köster *et al.*, *Neuroscience Letters*, **252** : 69-71 (1998)).

Taken together, these data suggest that the protein of SEQ ID NO: 214 may play a role in nervous system development and function. Thus, this protein may be useful in diagnosing and/or treating cancer and/or brain disorders, including neurodegenerative disorders such as Alzheimer's and Parkinson's diseases.

Protein of SEQ ID NO: 225 (internal designation 78-8-3-E6-CL0 1)

The protein of SEQ ID NO: 225 encoded by the extended cDNA SEQ ID NO: 124 and expressed in adult prostate belong to the phosphatidylethanolainin-binding protein from which it exhibits the characteristic PROSITE signature from positions 90 to 112 (see table VIII). Proteins from this widespread family, from nematodes to fly, yeast, rodent and primate species, bind hydrophobic ligands such as phospholipids and nucleotides. They are mostly expressed in brain and in testis and are thought to play a role in cell growth and/or maturation, in regulation of the sperm maturation, motility and in membrane remodeling. They may act either through signal transduction or through oxidoreduction reactions (for a review see Schoentgen and Jollès, *FEBS Letters*, **369** : 22-26 (1995)).

Taken together, these data suggest that the protein of SEQ ID NO: 225 may play a role in cell. Thus, these growth, maturation and in membrane remodeling and/or may be related to male fertility. Thus, this protein may be useful in diagnosing and/or treating cancer, neurodegenerative diseases, and/of, disorders related to male fertility and sterility.

Protein of SEQ ID NO: 153 (internal designation 33-10-4-H2-CL2 2)

The protein of SEQ ID NO: 153 encoded by the extended cDNA SEQ ID NO: 52 and expressed in brain exhibits homology to different integral membrane proteins. These membrane proteins include the nematode protein SRE-2 (Swissprot accession number Q09273) that belongs to the multigene SRE family of *C. elegans* receptor-like proteins and a family of tricarboxylate carriers conserved between flies and mammals. One member of this matched family is the rat tricarboxylate carrier (Genbank accession number S70011), an anion transporter localized in the inner membrane of mitochondria and involved in the biosynthesis of fatty acids and cholesterol. The protein of the invention contains a short transmembrane segments from positions 5 to 25 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, **10** :685-686 (1994)).

Taken together, these data suggest that the protein of SEQ ID NO: 153 may play a role in signal transduction and/or in molecule transport. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases, immune disorders, cardiovascular disorders, hypertension, renal injury and repair and septic shock

#### Protein of SEQ ID NO: 213 (internal designation 33-106-2-F10-CL1 3)

The protein of SEQ ID NO: 213 encoded by the extended cDNA SEQ ID NO: 112 and expressed in brain exhibits homology with part of the tRNA pseudouridine 55 synthase found in *Escherichia Coli* (Swissprot accession number P09171). This bacterial protein belongs to the NAP57/CBF5/TRUB family of nucleolar proteins found in bacteria, yeasts and mammals involved in rRNA or tRNA biosynthesis, ribosomal subunit assembly and/or centromere/microtubule binding.

Taken together, these data suggest that the protein of SEQ ID NO: 213 may play a role in rRNA or tRNA biogenesis and function. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, hearing loss or disorders linked to chromosomal instability such as dyskeratosis.

Protein of SEQ ED NO: 240 (internal designation 78-21-3-G7-CL2 1)

The protein of SEQ ID NO: 240 encoded by the extended cDNA SEQ ID NO: 139 and expressed in brain exhibits homology with a family of eukaryotic cell surface antigens containing 4 transmembrane domains. The PROSITE signature for this family is conserved in the protein of the invention except for a substitution of an alanine residue in place of any of the following hydrophobic residues : leucine, valine, isoleucine or methionine (positions 21 to 36).

The protein of the invention contains three short transmembrane segments from positions 6 to 26, 32 to 52 and from 56 to 76 as predicted by the software TopPred II (Claros and von Heijne, *CABIOS applic. Notes*, **10** : 685-686 (1994)). These transmembrane domains match the last three transmembrane domains of the matched protein family.

Taken together, these data suggest that the protein of SEQ ID NO: 240 may play a role in immunological and/or inflammatory responses, probably as a cell surface antigen. Thus, this protein or part therein, may be useful in diagnosing and treating several disorders including, but not limited to, cancer, immunological, haematological and/or inflammatory disorders. It may also be useful in modulating the immune and inflammatory responses to infectious agents and/or to suppress graft rejection.

Protein of SEQ ID NO: 239 (internal designation 76-30-3-B7-CL1 1)

The protein of SEQ ID NO: 239 encoded by the extended cDNA SEQ ID NO: 138 exhibits homology with a conserved region in a family of  $\text{Na}^+/\text{H}^+$  exchanger conserved in yeast, nematode and mammals. These cation/proton exchangers are integral membrane proteins with 5 transmembrane segments involved in intracellular pH regulation, maintenance of cell volume, reabsorption of sodium across specialized epithelia, vectorial transport and are also thought to play a role in signal transduction and especially in the induction of cell proliferation and in the induction of apoptosis.

The protein of invention contains four short transmembrane segments from positions 21 to 41, 48 to 68 and from 131 to 151 as predicted by the software TopPred II

(Claros and von Heijne, *CABIOS applic. Notes*, **10** : 685-686 (1994)). The third and fourth transmembrane domains match the fourth and fifth transmembrane segments of the matched family of proteins.

Taken together, these data suggest that the protein of SEQ ID NO: 239 may play a role in membrane permeability and/or in signal transduction. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases, cardiovascular disorders, hypertension, renal injury and repair, septic shock as well as disorders of membrane permeability such as diarrhea.

Protein of SEQ ID NO: 200 (internal designation 77-16-4-G3-CL1 3)

The protein of SEQ ID NO: 200 encoded by the extended cDNA SEQ ED NO: 99 and expressed in brain exhibits extensive homology to the N-terminus of cell division cycle protein 23 (Genbank accession number AF053977) and also to a lesser extent to its homologue in *Saccharomyces cerevisiae*. The matched protein is required for chromosome segregation and is part of the anaphase-promoting complex necessary for cell cycle progression to mitosis.

Taken together, these data suggest that the protein of SEQ ID NO: 200 may play a role in cellular mitosis. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer and leukemia.

Protein of SEQ ID NO: 230 (internal designation 33-61-2-F6-CL0 2)

The protein of SEQ ID NO: 230 encoded by the extended cDNA SEQ ID NO: 129 exhibits extensive homology to the C-terminus of the eta subunit of T-complex polypeptide 1 conserved from yeasts to mammals, and even complete identity with the last 54 amino acid residues of the human protein (Genbank accession number AF026292). The matched protein is a chaperonin which assists the folding of actins and tubulins in eukaryotic cells upon ATP hydrolysis.

Taken together, these data suggest that the protein of SEQ ID NO: 230 may play a role in the folding, transport, assembly and degradation of proteins. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, cardiovascular disorders, immune disorders, neurodegenerative disorders, osteoporosis and arthritis.

Protein of SEQ ED NO: 167 (internal designation 47-14-1-C3-CL0 5)

The protein of SEQ ID NO: 167 encoded by the extended cDNA SEQ ID NO: 66 exhibits homology to a monkey pepsinogen A-4 precursor (Swissprot accession number P27678) and to related members of the aspartyl protease family. The matched protein belongs to a family of widely distributed proteolytic enzymes known to exist in vertebrate, fungi, plants, retroviruses and some plant viruses.

Taken together, these data suggest that the protein of SEQ ID NO: 167 may play a role in the degradation of proteins. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, autoimmune disorders and immune disorders due to dysfunction of antigen presentation.

Protein of SEQ ID NO: 179 (internal designation 51-15-4-A12-CL11 3)

The protein of SEQ ID NO: 179 encoded by the extended cDNA SEQ ID NO: 78 found in testis exhibits homology to part of mammalian colipase precursors. Colipases are secreted cofactors for pancreatic lipases that allow the lipase to anchor at the water-lipid interface. Colipase plays a crucial role in the intestinal digestion and absorption of dietary fats. The 5 cysteines characteristic for this protein family are conserved in the protein of the invention although the colipase PROSITE signature is not.

Taken together, these data suggest that the protein of SEQ ED NO: 179 may play a role in the lipid metabolism and/or in male fertility. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, hyperlipidemia, hypercholesterolemia, atherosclerosis, cardiovascular disorders such

as coronary heart disease, and neurodegenerative disorders such as Alzheimer's disease or dementia, and disorders linked to male fertility.

Protein of SEQ ID NO: 227 (internal designation 33-11-1-B11-CL1 2)

5           The protein of SEQ ID NO: 227 encoded by the extended cDNA SEQ ID NO: 126 exhibits extensive homology to the ATP binding region of a whole family of serine/threonine protein kinases belonging to the CDC2/CDC28 subfamily. The PROSITE signature characteristic for this domain is present in the protein of the invention from positions 10 to 34.

10           Taken together, these data suggest that the protein of SEQ ED NO: 158 may bind ATP, and even be a protein kinase. Thus, this protein may be useful in diagnosing and/or treating several types of disorders including, but not limited to, cancer, neurodegenerative diseases, cardiovascular disorders, hypertension, renal injury and repair and septic shock.

15           As discussed above, the extended cDNAs of the present invention or portions thereof can be used for various purposes. The polynucleotides can be used to express recombinant protein for analysis, characterization or therapeutic use; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in disease states); as molecular weight markers on Southern gels; as chromosome markers or tags (when labeled) to  
20           identify chromosomes or to map related gene positions; to compare with endogenous DNA sequences in patients to identify potential genetic disorders; as probes to hybridize and thus discover novel, related DNA sequences; as a source of information to derive PCR primers for genetic fingerprinting; for selecting and making oligomers for attachment to a  
25           "gene chip" or other support, including for examination for expression patterns; to raise anti-protein antibodies using DNA immunization techniques; and as an antigen to raise anti-DNA antibodies or elicit another immune response. Where the polynucleotide encodes a protein which binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the polynucleotide can also be used in



interaction trap assays (such as, for example, that described in Gyuris et al., Cell 75:791-803 (1993)) to identify polynucleotides encoding the other protein with which binding occurs or to identify inhibitors of the binding interaction.

5 The proteins or polypeptides provided by the present invention can similarly be used in assays to determine biological activity, including in a panel of multiple proteins for high-throughput screening; to raise antibodies or to elicit another immune response; as a reagent (including the labeled reagent) in assays designed to quantitatively determine levels of the protein (or its receptor) in biological fluids; as markers for tissues in which the corresponding protein is preferentially expressed (either constitutively or at a particular stage of tissue differentiation or development or in a disease state); and, of course, to isolate correlative receptors or ligands. Where the protein binds or potentially binds to another protein (such as, for example, in a receptor-ligand interaction), the protein can be used to identify the other protein with which binding occurs or to identify inhibitors of the binding interaction. Proteins involved in these binding interactions can also be used to screen for peptide or small molecule inhibitors or agonists of the binding interaction.

Any or all of these research utilities are capable of being developed into reagent grade or kit format for commercialization as research products.

Methods for performing the uses listed above are well known to those skilled in the art. References disclosing such methods include without limitation "Molecular Cloning; A Laboratory Manual", 2d ed., Cole Spring Harbor Laboratory Press, Sambrook, J., E.F. Fritsch and T. Maniatis eds., 1989, and "Methods in Enzymology; Guide to Molecular Cloning Techniques", Academic Press, Berger, S.L. and A.R. Kimmel eds., 1987.

25 Polynucleotides and proteins of the present invention can also be used as nutritional sources or supplements. Such uses include without limitation use as a protein or amino acid supplement, use as a carbon source, use as a nitrogen source and use as a source of carbohydrate. In such cases the protein or polynucleotide of the invention can be added to the feed of a particular organism or can be administered as a separate solid or liquid preparation, such as in the form of powder, pills, solutions, suspensions or capsules.

In the case of microorganisms, the protein or polynucleotide of the invention can be added to the medium in or on which the microorganism is cultured.

5 Although this invention has been described in terms of certain preferred embodiments, other embodiments which will be apparent to those of ordinary skill in the art in view of the disclosure herein are also within the scope of this invention. Accordingly, the scope of the invention is intended to be defined only by reference to the appended claims. All documents cited herein are incorporated herein by reference in their entirety.

**TABLE I**

SEQ ID NO. in Present application	Provisional Application Disclosing Sequence	SEQ ID NO. in provisional application
40	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	51
41	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	72
42	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	52
43	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	78
44	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	73
45	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	41
46	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	67
47	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	82
48	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	80
49	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	81
50	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	53
51	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	54
52	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	195
53	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	44
54	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	46
55	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	68
56	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	48

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57	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	55
58	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	49
59	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	50
60	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	97
61	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	51
62	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	69
63	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	49
64	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	199
65	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	53
66	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	57
67	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	54
68	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	55
69	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	58
70	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	59
71	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	60
72	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	112
73	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	52
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77	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	75
78	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	61
79	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	61
80	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	130
81	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	65
82	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	54
83	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	78
84	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	63
85	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	65
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96	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	63
97	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	52
98	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	62
99	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	176
100	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	63
101	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	187
102	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	190
103	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	83
104	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	180
105	U.S. Provisional Patent Application Serial No. 60/096,116, filed Aug. 10, 1998	64
106	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	69
107	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	40
108	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	77
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113	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	46
114	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	47
115	U.S. Provisional Patent Application Serial No. 60/066,677, filed Nov. 13, 1997	53
116	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	58
117	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	74
118	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	71
119	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	145
120	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	67
121	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	58
122	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	72
123	U.S. Provisional Patent Application Serial No. 60/074,121, filed Feb. 9, 1998	73
124	U.S. Provisional Patent Application Serial No. 60/081,563, filed Apr. 13, 1998	70
125	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	40
126	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	44
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133	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	57
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136	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	64
137	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	65
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290	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	123
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312	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	145
313	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	146
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328	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	161
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330	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	163
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334	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	167
335	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	168
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343	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	176
344	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	177
345	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	178
346	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	179
347	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	180
348	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	181
349	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	182
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363	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	196
364	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	197
365	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	1998
366	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	199
367	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	200
368	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	201
369	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	202
370	U.S. Provisional Patent Application Serial No. 60/069,957, filed Dec. 17, 1997	203
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**TABLE II : Parameters used for each step of EST analysis**

Step	Search Characteristics			Selection Characteristics	
	Program	Strand	Parameters	Identity (%)	Length (bp)
Miscellaneous	Blastn	both	S=61 X=16	90	17
tRNA	Fasta	both	-	80	60
rRNA	Blastn	both	S=108	80	40
mtRNA	Blastn	both	S=108	80	40
Procaryotic	Blastn	both	S=144	90	40
Fungal	Blastn	both	S=144	90	40
Alu	fasta*	both	-	70	40
L1	Blastn	both	S=72	70	40
Repeats	Blastn	both	S=72	70	40
Promoters	Blastn	top	S=54 X=16	90	15†
Vertebrate	fasta*	both	S=108	90	30
ESTs	Blatsn	both	S=108 X=16	90	30
Proteins	blastx◇	top	E=0.001	-	-

\* use "Quick Fast" Database Scanner

† alignment further constrained to begin closer than 10bp to EST\5' end

◇ using BLOSUM62 substitution matrix

5

**TABLE III: Parameters used for each step of extended cDNA analysis**

Step	Search characteristics		Selection characteristics			
	Program	Strand	Parameters	Identity (%)	Length (bp)	Comments
miscellaneous*	FASTA	both	-	90	15	
tRNA <sup>s</sup>	FASTA	both	-	80	90	
rRNA <sup>s</sup>	BLASTN	both	S=108	80	40	
mtRNA <sup>s</sup>	BLASTN	both	S=108	80	40	
Procaryotic <sup>s</sup>	BLASTN	both	S=144	90	40	
Fungal*	BLASTN	both	S=144	90	40	
Alu*	BLASTN	both	S=72	70	40	max 5 matches, masking
LI <sup>s</sup>	BLASTN	both	S=72	70	40	max 5 matches, masking
Repeats <sup>s</sup>	BLASTN	both	S=72	70	40	masking
PolyA	BLAST2N	top	W=6,S=10,E=1000	90	8	in the last 20 nucleotides
Polyadenylation signal	-	top	AATAAA allowing 1 mismatch			in the 50 nucleotides preceding the 5' end of the polyA
Vertibrate*	BLASTN then FASTA	both	-	90 then 70	30	first BLASTN and then FASTA on matching sequences
ESTs*	BLAST2N	both	-	90	30	
Geneseq	BLASTN	both	W=8, B=10	90	30	
ORF	BLASTP	top	W=8, B=10	-	-	on ORF proteins, max 10 matches
Proteins*	BLASTX	top	E=0.001	70	30	

<sup>s</sup> steps common to EST analysis and using the same algorithms and parameters

5 \* steps also used in EST analysis but with different algorithms and/or parameters

TABLE IV

Id	FCS Location	SigPep Location	Mature Polypeptide Location	Stop Codon Location	PolyA Signal Location	PolyA Site Location
40	7 through 471	7 through 99	100 through 471	472	537 through 542	554 through 568
41	168 through 332	-	168 through 332	333	557 through 562	-
42	51 through 251	51 through 110	111 through 251	252	849 through 854	882 through 895
43	20 through 613	20 through 82	83 through 613	614	-	-
44	12 through 416	12 through 86	87 through 416	417	425 through 430	445 through 458
45	276 through 1040	276 through 485	486 through 1040	1041	-	2024 through 2036
46	443 through 619	443 through 589	590 through 619	620	-	1267 through 1276
47	206 through 747	-	206 through 747	-	-	-
48	36 through 521	36 through 104	105 through 521	522	528 through 533	548 through 561
49	36 through 395	36 through 104	105 through 395	396	599 through 604	619 through 632
50	21 through 41	-	21 through 41	42	328 through 333	357 through 370
51	35 through 631	35 through 160	161 through 631	632	901 through 906	979 through 994
52	271 through 399	-	271 through 399	400	-	-
53	103 through 252	103 through 213	214 through 252	253	-	588 through 597
54	2 through 460	-	2 through 460	461	713 through 718	735 through 748
55	31 through 231	-	31 through 231	232	769 through 774	690 through 703
56	305 through 565	-	305 through 565	566	694 through 699	713 through 725
57	124 through 873	124 through 378	379 through 873	874	1673 through 1678	1694 through 1705
58	135 through 206	-	135 through 206	207	850 through 855	1056 through 1069
59	135 through 818	-	135 through 818	819	909 through 914	1071 through 1084
60	33 through 290	33 through 92	93 through 290	291	-	-
61	485 through 616	-	485 through 616	617	-	669 through 682
62	54 through 995	54 through 227	228 through 995	996	1130 through 1135	1181 through 1191
63	657 through 923	657 through 896	897 through 923	924	957 through 962	974 through 1008
64	18 through 311	18 through 62	63 through 311	312	-	-
65	151 through 426	151 through 258	259 through 426	427	505 through 510	527 through 538
66	10 through 1062	10 through 57	58 through 1062	1063	1710 through 1715	1735 through 1747
67	78 through 491	78 through 218	219 through 491	492	1652 through 1657	1673 through 1686
68	69 through 371	69 through 287	288 through 371	372	510 through 515	530 through 542
69	2 through 757	2 through 205	206 through 757	758	-	1160 through 1174
70	2 through 1051	2 through 205	206 through 1051	1052	1248 through 1253	1272 through 1285
71	2 through 1171	2 through 205	206 through 1171	1172	1368 through 1373	1386 through 1398
72	42 through 611	42 through 287	288 through 611	612	787 through 792	808 through 821
73	62 through 916	62 through 757	758 through 916	-	-	904 through 916
74	62 through 520	-	62 through 520	521	1124 through 1129	1141 through 1153

CONT. TABLE IV

75	21 through 167	-	21 through 167	168	-	-
76	22 through 318	22 through 93	94 through 318	319	497 through 502	516 through 526
77	8 through 292	8 through 118	119 through 292	293	317 through 322	339 through 352
78	16 through 378	16 through 84	85 through 378	379	502 through 507	522 through 542
79	57 through 233	-	57 through 233	-	-	-
80	83 through 340	83 through 124	125 through 340	341	573 through 578	607 through 660
81	47 through 541	47 through 220	221 through 541	542	-	597 through 605
82	46 through 285	46 through 150	151 through 285	286	364 through 369	385 through 396
83	22 through 240	22 through 84	85 through 240	241	397 through 402	421 through 432
84	89 through 382	-	89 through 382	383	-	408 through 420
85	80 through 415	80 through 142	143 through 415	416	471 through 476	488 through 501
86	152 through 361	152 through 283	284 through 361	362	-	-
87	32 through 307	32 through 70	71 through 307	308	1240 through 1245	1261 through 1272
88	114 through 734	114 through 239	240 through 734	735	768 through 773	793 through 804
89	199 through 802	-	199 through 802	-	780 through 785	791 through 802
90	38 through 1174	38 through 148	149 through 1174	1175	1452 through 1457	1478 through 1490
91	26 through 361	-	26 through 361	-	-	350 through 361
92	3 through 131	-	3 through 131	132	-	591 through 605
93	33 through 185	33 through 80	81 through 185	186	570 through 575	586 through 591
94	184 through 915	184 through 237	238 through 915	916	1119 through 1124	1139 through 1150
95	58 through 1116	58 through 159	160 through 1116	1117	1486 through 1491	1504 through 1513
96	327 through 417	-	327 through 417	-	-	404 through 417
97	63 through 398	63 through 206	207 through 398	399	-	-
98	2 through 163	-	2 through 163	164	488 through 493	511 through 522
99	13 through 465	13 through 75	76 through 465	466	-	-
100	20 through 703	20 through 94	95 through 703	704	1000 through 1005	1023 through 1041
101	103 through 294	103 through 243	244 through 294	295	-	-
102	81 through 518	81 through 173	174 through 518	519	-	-
103	66 through 326	-	66 through 326	327	1066 through 1071	1087 through 1098
104	170 through 289	170 through 250	251 through 289	290	-	-
105	36 through 497	-	36 through 497	498	650 through 655	663 through 685
106	18 through 320	-	18 through 320	321	539 through 544	542 through 554
107	71 through 1438	71 through 136	137 through 1438	1439	1644 through 1649	1665 through 1678
108	25 through 318	25 through 75	76 through 318	319	452 through 457	482 through 494
109	84 through 332	84 through 170	171 through 332	333	-	702 through 714
110	32 through 718	32 through 100	101 through 718	719	770 through 775	793 through 805
111	26 through 481	26 through 88	89 through 481	482	755 through 760	775 through 787

CONT. TABLE IV

112	26 through 562	26 through 187	188 through 562	563	-	-
113	4 through 810	4 through 279	280 through 810	811	858 through 863	881 through 893
114	55 through 459	55 through 120	121 through 459	460	1444 through 1449	1462 through 1475
115	48 through 248	48 through 161	162 through 248	249	283 through 288	308 through 321
116	25 through 399	25 through 186	187 through 399	400	-	-
117	10 through 1137	10 through 72	73 through 1137	1138	1144 through 1149	1162 through 1173
118	72 through 704	72 through 161	162 through 704	705	772 through 777	-
119	44 through 505	44 through 223	224 through 505	506	-	-
120	25 through 393	25 through 150	151 through 393	394	734 through 739	757 through 770
121	58 through 1095	58 through 114	115 through 1095	1096	-	1202 through 1213
122	31 through 660	31 through 90	91 through 660	661	1288 through 1293	1307 through 1318
123	31 through 582	31 through 90	91 through 582	583	816 through 821	840 through 853
124	15 through 695	15 through 80	81 through 695	696	795 through 800	814 through 826
125	74 through 295	74 through 196	197 through 295	296	545 through 550	561 through 571
126	440 through 659	-	440 through 659	-	601 through 606	-
127	38 through 283	38 through 85	86 through 283	284	257 through 262	-
128	121 through 477	121 through 288	289 through 477	-	-	-
129	2 through 163	-	2 through 163	164	292 through 297	310 through 323
130	46 through 675	46 through 87	88 through 675	676	1364 through 1369	1383 through 1392
131	62 through 385	-	62 through 385	386	974 through 979	987 through 999
132	422 through 550	422 through 475	476 through 550	551	-	714 through 725
133	124 through 231	-	124 through 231	232	-	387 through 400
134	131 through 1053	131 through 169	170 through 1053	-	1019 through 1024	-
135	86 through 403	86 through 181	182 through 403	404	1097 through 1102	1117 through 1128
136	37 through 162	37 through 93	94 through 162	163	224 through 229	243 through 254
137	31 through 381	31 through 90	91 through 381	382	-	875 through 886
138	46 through 579	46 through 156	157 through 579	580	-	-
139	92 through 471	92 through 172	173 through 471	-	454 through 459	458 through 471
140	154 through 675	154 through 498	499 through 675	676	819 through 824	838 through 849
242	18 through 173	18 through 77	78 through 173	174	864 through 869	882 through 893
243	17 through 595	17 through 85	86 through 595	596	820 through 825	840 through 851
244	89 through 334	89 through 130	131 through 334	335	462 through 467	484 through 495
245	21 through 614	21 through 83	84 through 614	615	849 through 854	873 through 884
246	94 through 573	94 through 258	259 through 573	574	862 through 867	886 through 897
247	74 through 397	74 through 127	128 through 397	398	472 through 477	507 through 518
248	51 through 242	51 through 116	117 through 242	243	319 through 324	339 through 350
249	111 through 191	111 through 155	156 through 191	192	965 through 970	986 through 996

CONT. TABLE IV

250	45 through 602	45 through 107	108 through 602	603	828 through 833	850 through 860
251	24 through 560	24 through 101	102 through 560	561	563 through 568	583 through 593
252	109 through 558	109 through 273	274 through 558	559	-	1104 through 1114
253	128 through 835	128 through 220	221 through 835	836	1145 through 1150	1170 through 1181
254	59 through 505	59 through 358	359 through 505	506	1042 through 1047	1062 through 1073
255	1 through 207	1 through 147	148 through 207	208	784 through 789	807 through 818
256	12 through 734	12 through 101	102 through 734	735	914 through 919	961 through 971
257	378 through 518	378 through 467	468 through 518	519	607 through 612	628 through 640
258	110 through 304	110 through 193	194 through 304	305	708 through 713	732 through 743
259	201 through 419	201 through 272	273 through 419	420	601 through 606	627 through 637
260	123 through 302	123 through 176	177 through 302	303	1279 through 1284	1301 through 1312
261	98 through 673	98 through 376	377 through 673	674	-	1025 through 1035
262	17 through 463	17 through 232	233 through 463	464	657 through 662	684 through 696
263	263 through 481	263 through 322	323 through 481	482	-	858 through 868
264	42 through 299	42 through 101	102 through 299	300	-	762 through 775
265	198 through 431	198 through 260	261 through 431	432	-	1064 through 1074
266	279 through 473	279 through 362	363 through 473	474	944 through 949	970 through 981
267	12 through 644	12 through 92	93 through 644	645	1002 through 1007	1020 through 1031
268	91 through 459	91 through 330	331 through 459	460	-	1271 through 1281
269	70 through 327	70 through 147	148 through 327	328	1741 through 1746	1763 through 1774
270	12 through 497	12 through 104	105 through 497	498	935 through 940	955 through 967
271	90 through 383	90 through 200	201 through 383	384	609 through 614	632 through 643
272	332 through 541	332 through 376	377 through 541	542	739 through 744	761 through 773
273	43 through 222	43 through 177	178 through 222	223	530 through 535	555 through 566
274	115 through 231	115 through 180	181 through 231	232	419 through 424	445 through 455
275	232 through 384	232 through 300	301 through 384	385	650 through 655	662 through 673
276	143 through 427	143 through 286	287 through 427	428	606 through 611	628 through 639
277	284 through 463	284 through 379	380 through 463	464	-	762 through 772
278	162 through 671	162 through 398	399 through 671	672	805 through 810	830 through 840
279	63 through 632	63 through 308	309 through 632	633	808 through 813	829 through 840
280	21 through 362	21 through 200	201 through 362	363	821 through 826	838 through 849
281	21 through 503	21 through 344	345 through 503	504	1305 through 1310	1330 through 1341
282	1 through 201	1 through 63	64 through 201	202	637 through 642	660 through 671
283	39 through 1034	39 through 134	135 through 1034	1035	1566 through 1571	1587 through 1597
284	69 through 263	69 through 125	126 through 263	264	1173 through 1178	1196 through 1205
285	115 through 285	115 through 204	205 through 285	286	505 through 510	525 through 536
286	90 through 344	90 through 140	141 through 344	345	500 through 505	515 through 527

CONT. TABLE IV

287	57 through 311	57 through 107	108 through 311	312	467 through 472	482 through 493
288	96 through 302	96 through 182	183 through 302	303	-	501 through 514
289	161 through 526	161 through 328	329 through 526	527	-	799 through 811
290	210 through 332	210 through 299	300 through 332	333	594 through 599	613 through 625
291	212 through 361	212 through 319	320 through 361	362	650 through 655	673 through 684
292	75 through 482	75 through 128	129 through 482	483	595 through 600	618 through 627
293	50 through 631	50 through 244	245 through 631	632	777 through 782	801 through 812
294	154 through 576	154 through 360	361 through 576	577	737 through 742	763 through 775
295	154 through 897	154 through 360	361 through 897	898	1017 through 1022	1044 through 1054
296	146 through 292	146 through 253	254 through 292	293	395 through 400	433 through 444
297	126 through 383	126 through 167	168 through 383	384	726 through 731	743 through 754
298	66 through 497	66 through 239	240 through 497	498	594 through 599	618 through 629
299	49 through 411	49 through 96	97 through 411	412	732 through 737	750 through 763
300	49 through 534	49 through 96	97 through 534	535	593 through 598	612 through 623
301	86 through 415	86 through 145	146 through 415	416	540 through 545	560 through 571
302	56 through 268	56 through 100	101 through 268	269	584 through 589	601 through 612
303	32 through 328	32 through 103	104 through 328	329	508 through 513	528 through 539
304	21 through 527	21 through 95	96 through 527	528	921 through 926	953 through 963
305	147 through 647	147 through 374	375 through 647	648	-	668 through 681
306	262 through 471	262 through 306	307 through 471	472	663 through 668	682 through 693
307	74 through 1216	74 through 172	173 through 1216	1217	1627 through 1632	1640 through 1652
308	48 through 164	48 through 89	90 through 164	165	482 through 487	505 through 517
309	185 through 334	185 through 295	296 through 334	335	355 through 360	392 through 405
310	195 through 347	195 through 272	273 through 347	348	1037 through 1042	1071 through 1082
311	90 through 815	90 through 179	180 through 815	816	883 through 888	905 through 916
312	52 through 513	52 through 231	232 through 513	514	553 through 558	572 through 583
313	172 through 438	172 through 354	355 through 438	439	682 through 687	685 through 697
314	148 through 366	148 through 225	226 through 366	367	770 through 775	792 through 803
315	175 through 336	175 through 276	277 through 336	337	-	812 through 823
316	191 through 553	191 through 304	305 through 553	554	766 through 771	804 through 817
317	106 through 603	106 through 216	217 through 603	604	-	1102 through 1112
318	47 through 586	47 through 124	125 through 586	587	1583 through 1588	1614 through 1623
319	99 through 371	99 through 290	291 through 371	372	491 through 496	513 through 524
320	44 through 814	44 through 112	113 through 814	815	-	978 through 989
321	3 through 581	3 through 182	183 through 581	582	-	1006 through 1016
322	107 through 427	107 through 190	191 through 427	428	499 through 504	516 through 529
323	45 through 407	45 through 83	84 through 407	408	1008 through 1013	1032 through 1042

CONT. TABLE IV

324	201 through 332	201 through 251	252 through 332	333	-	869 through 880
325	217 through 543	217 through 255	256 through 543	544	-	1206 through 1217
326	18 through 446	18 through 140	141 through 446	447	930 through 935	948 through 959
327	29 through 724	29 through 118	119 through 724	725	886 through 891	910 through 920
328	404 through 586	404 through 466	467 through 586	587	1304 through 1309	1334 through 1344
329	331 through 432	331 through 387	388 through 432	433	548 through 553	573 through 585
330	59 through 703	59 through 220	221 through 703	704	886 through 891	903 through 914
331	672 through 752	672 through 722	723 through 752	753	-	1150 through 1161
332	57 through 311	57 through 128	129 through 311	312	332 through 337	351 through 363
333	80 through 232	80 through 127	128 through 232	233	617 through 622	634 through 645
334	91 through 291	91 through 219	220 through 291	292	367 through 372	389 through 400
335	196 through 384	196 through 240	241 through 384	385	461 through 466	485 through 496
336	54 through 590	54 through 227	228 through 590	591	-	955 through 965
337	133 through 846	133 through 345	346 through 846	847	-	890 through 901
338	138 through 671	138 through 248	249 through 671	672	1319 through 1324	1338 through 1347
339	124 through 411	124 through 186	187 through 411	412	948 through 953	971 through 983
340	372 through 494	372 through 443	444 through 494	495	708 through 713	732 through 745
341	112 through 450	112 through 192	193 through 450	451	1053 through 1058	1095 through 1106
342	117 through 866	117 through 170	171 through 866	867	1159 through 1164	1178 through 1190
343	13 through 465	13 through 75	76 through 465	466	1035 through 1040	1060 through 1070
344	2 through 718	2 through 76	77 through 718	719	1170 through 1175	1203 through 1213
345	86 through 709	86 through 361	362 through 709	710	943 through 948	963 through 973
346	63 through 320	63 through 179	180 through 320	321	771 through 776	799 through 810
347	299 through 418	299 through 379	380 through 418	419	739 through 744	762 through 771
348	186 through 380	186 through 233	234 through 380	381	383 through 388	396 through 409
349	69 through 458	69 through 233	234 through 458	459	564 through 569	602 through 613
350	12 through 638	12 through 263	264 through 638	639	951 through 956	975 through 985
351	282 through 389	282 through 332	333 through 389	390	1413 through 1418	1437 through 1447
352	208 through 339	208 through 294	295 through 339	340	-	1631 through 1641
353	69 through 557	69 through 224	225 through 557	558	849 through 854	870 through 883
354	134 through 325	134 through 274	275 through 325	326	-	718 through 729
355	78 through 731	78 through 227	228 through 731	732	-	1002 through 1013
356	46 through 693	46 through 90	91 through 693	694	937 through 942	962 through 973
357	126 through 527	126 through 182	183 through 527	528	834 through 839	856 through 867
358	66 through 320	66 through 113	114 through 320	321	490 through 495	508 through 519
359	73 through 948	73 through 159	160 through 948	949	-	1016 through 1028
360	69 through 434	69 through 236	237 through 434	435	419 through 424	441 through 452



CONT. TABLE IV

361	628 through 804	628 through 711	712 through 804	805	-	864 through 875
362	70 through 366	70 through 108	109 through 366	367	496 through 501	521 through 531
363	70 through 366	70 through 108	109 through 366	367	-	1233 through 1244
364	111 through 434	111 through 185	186 through 434	435	-	618 through 631
365	19 through 567	19 through 63	64 through 567	568	749 through 754	771 through 781
366	19 through 312	19 through 63	64 through 312	313	896 through 901	921 through 931
367	64 through 612	64 through 234	235 through 612	613	-	839 through 849
368	39 through 458	39 through 80	81 through 458	459	613 through 618	633 through 644
369	9 through 185	9 through 50	51 through 185	186	-	906 through 918
370	14 through 316	14 through 121	122 through 316	317	442 through 447	458 through 471
371	70 through 1092	70 through 234	235 through 1092	1093	1475 through 1480	1493 through 1504
372	274 through 597	274 through 399	400 through 597	598	731 through 736	754 through 765
373	230 through 469	230 through 307	308 through 469	470	1004 through 1009	1027 through 1040
374	72 through 545	72 through 203	204 through 545	546	-	1151 through 1162
375	36 through 425	36 through 119	120 through 425	426	1215 through 1220	1240 through 1250
376	155 through 751	155 through 340	341 through 751	752	912 through 917	937 through 947
377	46 through 585	46 through 120	121 through 585	586	584 through 589	606 through 619

TABLE V

<b>Id</b>	<b>Full Length Polypeptide Location</b>	<b>Signal Peptide Location</b>	<b>Mature Polypeptide Location</b>
141	-31 through 124	-31 through -1	1 through 124
142	1 through 55	-	1 through 55
143	-20 through 47	-20 through -1	1 through 47
144	-21 through 177	-21 through -1	1 through 177
145	-25 through 110	-25 through -1	1 through 110
146	-70 through 185	-70 through -1	1 through 185
147	-49 through 10	-49 through -1	1 through 10
148	1 through 180	-	1 through 180
149	-23 through 139	-23 through -1	1 through 139
150	-23 through 97	-23 through -1	1 through 97
151	1 through 7	-	1 through 7
152	-42 through 157	-42 through -1	1 through 157
153	1 through 43	-	1 through 43
154	-37 through 13	-37 through -1	1 through 13
155	1 through 153	-	1 through 153
156	1 through 67	-	1 through 67
157	1 through 87	-	1 through 87
158	-85 through 165	-85 through -1	1 through 165
159	1 through 24	-	1 through 24
160	1 through 228	-	1 through 228
161	-20 through 66	-20 through -1	1 through 66
162	1 through 44	-	1 through 44
163	-58 through 256	-58 through -1	1 through 256
164	-80 through 9	-80 through -1	1 through 9
165	-15 through 83	-15 through -1	1 through 83
166	-36 through 56	-36 through -1	1 through 56
167	-16 through 335	-16 through -1	1 through 335
168	-47 through 91	-47 through -1	1 through 91
169	-73 through 28	-73 through -1	1 through 28
170	-68 through 184	-68 through -1	1 through 184
171	-68 through 282	-68 through -1	1 through 282
172	-68 through 322	-68 through -1	1 through 322
173	-82 through 108	-82 through -1	1 through 108
174	-232 through 53	-232 through -1	1 through 53
175	1 through 153	-	1 through 153
176	1 through 49	-	1 through 49
177	-24 through 75	-24 through -1	1 through 75
178	-37 through 58	-37 through -1	1 through 58
179	-23 through 98	-23 through -1	1 through 98
180	1 through 59	-	1 through 59
181	-14 through 72	-14 through -1	1 through 72
182	-58 through 107	-58 through -1	1 through 107
183	-35 through 45	-35 through -1	1 through 45

CONT. TABLE V

184	-21 through 52	-21 through -1	1 through 52
185	1 through 98	-	1 through 98
186	-21 through 91	-21 through -1	1 through 91
187	-44 through 26	-44 through -1	1 through 26
188	-13 through 79	-13 through -1	1 through 79
189	-42 through 165	-42 through -1	1 through 165
190	1 through 201	-	1 through 201
191	-37 through 342	-37 through -1	1 through 342
192	1 through 112	-	1 through 112
193	1 through 43	-	1 through 43
194	-16 through 35	-16 through -1	1 through 35
195	-18 through 226	-18 through -1	1 through 226
196	-34 through 319	-34 through -1	1 through 319
197	1 through 30	-	1 through 30
198	-48 through 64	-48 through -1	1 through 64
199	1 through 54	-	1 through 54
200	-21 through 130	-21 through -1	1 through 130
201	-25 through 203	-25 through -1	1 through 203
202	-47 through 17	-47 through -1	1 through 17
203	-31 through 115	-31 through -1	1 through 115
204	1 through 87	-	1 through 87
205	-27 through 13	-27 through -1	1 through 13
206	1 through 154	-	1 through 154
207	1 through 101	-	1 through 101
208	-22 through 434	-22 through -1	1 through 434
209	-17 through 81	-17 through -1	1 through 81
210	-29 through 54	-29 through -1	1 through 54
211	-23 through 206	-23 through -1	1 through 206
212	-21 through 131	-21 through -1	1 through 131
213	-54 through 125	-54 through -1	1 through 125
214	-92 through 177	-92 through -1	1 through 177
215	-22 through 113	-22 through -1	1 through 113
216	-38 through 29	-38 through -1	1 through 29
217	-54 through 71	-54 through -1	1 through 71
218	-21 through 355	-21 through -1	1 through 355
219	-30 through 181	-30 through -1	1 through 181
220	-60 through 94	-60 through -1	1 through 94
221	-42 through 81	-42 through -1	1 through 81
222	-19 through 327	-19 through -1	1 through 327
223	-20 through 190	-20 through -1	1 through 190
224	-20 through 164	-20 through -1	1 through 164
225	-22 through 205	-22 through -1	1 through 205
226	-41 through 33	-41 through -1	1 through 33
227	1 through 73	-	1 through 73
228	-16 through 66	-16 through -1	1 through 66
229	-56 through 63	-56 through -1	1 through 63

CONT. TABLE V

230	1 through 54	-	1 through 54
231	-14 through 196	-14 through -1	1 through 196
232	1 through 108	-	1 through 108
233	-18 through 25	-18 through -1	1 through 25
234	1 through 36	-	1 through 36
235	-13 through 294	-13 through -1	1 through 294
236	-32 through 74	-32 through -1	1 through 74
237	-19 through 23	-19 through -1	1 through 23
238	-20 through 97	-20 through -1	1 through 97
239	-37 through 141	-37 through -1	1 through 141
240	-27 through 99	-27 through -1	1 through 99
241	-115 through 59	-115 through -1	1 through 59
378	-20 through 32	-20 through -1	1 through 32
379	-23 through 170	-23 through -1	1 through 170
380	-14 through 68	-14 through -1	1 through 68
381	-21 through 177	-21 through -1	1 through 177
382	-55 through 105	-55 through -1	1 through 105
383	-18 through 90	-18 through -1	1 through 90
384	-22 through 42	-22 through -1	1 through 42
385	-15 through 12	-15 through -1	1 through 12
386	-21 through 165	-21 through -1	1 through 165
387	-26 through 153	-26 through -1	1 through 153
388	-55 through 95	-55 through -1	1 through 95
389	-31 through 205	-31 through -1	1 through 205
390	-100 through 49	-100 through -1	1 through 49
391	-49 through 20	-49 through -1	1 through 20
392	-30 through 211	-30 through -1	1 through 211
393	-30 through 17	-30 through -1	1 through 17
394	-28 through 37	-28 through -1	1 through 37
395	-24 through 49	-24 through -1	1 through 49
396	-18 through 42	-18 through -1	1 through 42
397	-93 through 99	-93 through -1	1 through 99
398	-72 through 77	-72 through -1	1 through 77
399	-20 through 53	-20 through -1	1 through 53
400	-20 through 66	-20 through -1	1 through 66
401	-21 through 57	-21 through -1	1 through 57
402	-28 through 37	-28 through -1	1 through 37
403	-27 through 184	-27 through -1	1 through 184
404	-80 through 43	-80 through -1	1 through 43
405	-26 through 60	-26 through -1	1 through 60
406	-31 through 131	-31 through -1	1 through 131
407	-37 through 61	-37 through -1	1 through 61
408	-15 through 55	-15 through -1	1 through 55
409	-45 through 15	-45 through -1	1 through 15
410	-22 through 17	-22 through -1	1 through 17
411	-23 through 28	-23 through -1	1 through 28

CONT. TABLE V

412	-48 through 47	-48 through -1	1 through 47
413	-32 through 28	-32 through -1	1 through 28
414	-79 through 91	-79 through -1	1 through 91
415	-82 through 108	-82 through -1	1 through 108
416	-60 through 54	-60 through -1	1 through 54
417	-108 through 53	-108 through -1	1 through 53
418	-21 through 46	-21 through -1	1 through 46
419	-32 through 300	-32 through -1	1 through 300
420	-19 through 46	-19 through -1	1 through 46
422	-30 through 27	-30 through -1	1 through 27
423	-17 through 68	-17 through -1	1 through 68
424	-17 through 68	-17 through -1	1 through 68
425	-29 through 40	-29 through -1	1 through 40
426	-56 through 66	-56 through -1	1 through 66
427	-30 through 11	-30 through -1	1 through 11
428	-36 through 14	-36 through -1	1 through 14
429	-18 through 118	-18 through -1	1 through 118
430	-65 through 129	-65 through -1	1 through 129
431	-69 through 72	-69 through -1	1 through 72
432	-69 through 179	-69 through -1	1 through 179
433	-36 through 13	-36 through -1	1 through 13
434	-14 through 72	-14 through -1	1 through 72
435	-58 through 86	-58 through -1	1 through 86
436	-16 through 105	-16 through -1	1 through 105
437	-16 through 146	-16 through -1	1 through 146
438	-20 through 90	-20 through -1	1 through 90
439	-15 through 56	-15 through -1	1 through 56
440	-24 through 75	-24 through -1	1 through 75
441	-25 through 144	-25 through -1	1 through 144
442	-76 through 91	-76 through -1	1 through 91
443	-15 through 55	-15 through -1	1 through 55
444	-33 through 348	-33 through -1	1 through 348
445	-14 through 25	-14 through -1	1 through 25
446	-37 through 13	-37 through -1	1 through 13
447	-26 through 25	-26 through -1	1 through 25
448	-30 through 212	-30 through -1	1 through 212
449	-60 through 94	-60 through -1	1 through 94
450	-61 through 28	-61 through -1	1 through 28
451	-26 through 47	-26 through -1	1 through 47
452	-34 through 20	-34 through -1	1 through 20
453	-38 through 83	-38 through -1	1 through 83
454	-37 through 129	-37 through -1	1 through 129
455	-26 through 154	-26 through -1	1 through 154
456	-64 through 27	-64 through -1	1 through 27
457	-23 through 234	-23 through -1	1 through 234
458	-60 through 133	-60 through -1	1 through 133

CONT. TABLE V

459	-28 through 79	-28 through -1	1 through 79
460	-13 through 108	-13 through -1	1 through 108
461	-17 through 27	-17 through -1	1 through 27
462	-13 through 96	-13 through -1	1 through 96
463	-41 through 102	-41 through -1	1 through 102
464	-30 through 202	-30 through -1	1 through 202
465	-21 through 40	-21 through -1	1 through 40
466	-19 through 15	-19 through -1	1 through 15
467	-54 through 161	-54 through -1	1 through 161
468	-17 through 10	-17 through -1	1 through 10
469	-24 through 61	-24 through -1	1 through 61
470	-16 through 35	-16 through -1	1 through 35
471	-43 through 24	-43 through -1	1 through 24
472	-15 through 48	-15 through -1	1 through 48
473	-58 through 121	-58 through -1	1 through 121
474	-71 through 167	-71 through -1	1 through 167
475	-37 through 141	-37 through -1	1 through 141
476	-21 through 75	-21 through -1	1 through 75
477	-24 through 17	-24 through -1	1 through 17
478	-27 through 86	-27 through -1	1 through 86
479	-18 through 232	-18 through -1	1 through 232
480	-21 through 130	-21 through -1	1 through 130
481	-25 through 214	-25 through -1	1 through 214
482	-92 through 116	-92 through -1	1 through 116
483	-39 through 47	-39 through -1	1 through 47
484	-27 through 13	-27 through -1	1 through 13
485	-16 through 49	-16 through -1	1 through 49
486	-55 through 75	-55 through -1	1 through 75
487	-84 through 125	-84 through -1	1 through 125
488	-17 through 19	-17 through -1	1 through 19
489	-29 through 15	-29 through -1	1 through 15
490	-52 through 111	-52 through -1	1 through 111
491	-47 through 17	-47 through -1	1 through 17
492	-50 through 168	-50 through -1	1 through 168
493	-15 through 201	-15 through -1	1 through 201
494	-19 through 115	-19 through -1	1 through 115
495	-16 through 69	-16 through -1	1 through 69
496	-29 through 263	-29 through -1	1 through 263
497	-56 through 66	-56 through -1	1 through 66
498	-28 through 31	-28 through -1	1 through 31
499	-13 through 86	-13 through -1	1 through 86
500	-13 through 86	-13 through -1	1 through 86
501	-25 through 83	-25 through -1	1 through 83
502	-15 through 168	-15 through -1	1 through 168
503	-15 through 83	-15 through -1	1 through 83
504	-57 through 126	-57 through -1	1 through 126

CONT. TABLE V

505	-14 through 126	-14 through -1	1 through 126
506	-14 through 45	-14 through -1	1 through 45
507	-36 through 65	-36 through -1	1 through 65
508	-55 through 286	-55 through -1	1 through 286
509	-42 through 66	-42 through -1	1 through 66
510	-26 through 54	-26 through -1	1 through 54
511	-44 through 114	-44 through -1	1 through 114
512	-28 through 102	-28 through -1	1 through 102
513	-62 through 137	-62 through -1	1 through 137
514	-25 through 155	-25 through -1	1 through 155

TABLE VI

<b>Id</b>	<b>Collection Refs</b>	<b>Deposit Name</b>
40	ATCC # 98921	SignalTag 121-144
41	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
42	ATCC # 98921	SignalTag 121-144
43	ATCC # 98920	SignalTag 67-90
44	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
45	ATCC # 98920	SignalTag 67-90
46	ATCC # 98923	SignalTag 44-66
47	ATCC # 98920	SignalTag 67-90
48	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
49	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
50	ATCC # 98921	SignalTag 121-144
51	ATCC # 98921	SignalTag 121-144
52	ATCC # 98920	SignalTag 67-90
53	ATCC # 98923	SignalTag 44-66
54	ATCC # 98920	SignalTag 67-90
55	ATCC # 98920	SignalTag 67-90
56	ATCC # 98920	SignalTag 67-90
57	ATCC # 98921	SignalTag 121-144
58	ATCC # 98920	SignalTag 67-90
59	ATCC # 98920	SignalTag 67-90
60	ATCC # 98920	SignalTag 67-90
61	ATCC # 98923	SignalTag 44-66
62	ATCC # 98923	SignalTag 44-66
63	ATCC # 98923	SignalTag 44-66
64	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
65	ATCC # 98923	SignalTag 44-66
66	ATCC # 98921	SignalTag 121-144
67	ATCC # 98920	SignalTag 67-90
68	ATCC # 98920	SignalTag 67-90



CONT. TABLE VI

69	ATCC # 98921	SignalTag 121-144
70	ATCC # 98921	SignalTag 121-144
71	ATCC # 98921	SignalTag 121-144
72	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
73	ATCC # 98923	SignalTag 44-66
74	ATCC # 98923	SignalTag 44-66
75	ATCC # 98920	SignalTag 67-90
76	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
77	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
78	ATCC # 98921	SignalTag 121-144
79	ATCC # 98923	SignalTag 44-66
80	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
81	ATCC # 98921	SignalTag 121-144
82	ATCC # 98920	SignalTag 67-90
83	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
84	ATCC # 98923	SignalTag 44-66
85	ATCC # 98923	SignalTag 44-66
86	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
87	ATCC # 98923	SignalTag 44-66
88	ATCC # 98923	SignalTag 44-66
89	ATCC # 98923	SignalTag 44-66
90	ATCC # 98923	SignalTag 44-66
91	ATCC # 98923	SignalTag 44-66
92	ATCC # 98920	SignalTag 67-90
93	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
94	ATCC # 98923	SignalTag 44-66
95	ATCC # 98923	SignalTag 44-66
96	ATCC # 98920	SignalTag 67-90
97	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
98	ATCC # 98921	SignalTag 121-144
99	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120

CONT. TABLE VI

100	ATCC # 98921	SignalTag 121-144
101	ATCC # 98920	SignalTag 67-90
102	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
103	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
104	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
105	ATCC # 98921	SignalTag 121-144
106	ATCC # 98920	SignalTag 67-90
107	ATCC # 98920	SignalTag 67-90
108	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
109	ATCC # 98923	SignalTag 44-66
110	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
111	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
112	ATCC # 98920	SignalTag 67-90
113	ATCC # 98920	SignalTag 67-90
114	ATCC # 98923	SignalTag 44-66
115	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
116	ATCC # 98920	SignalTag 67-90
117	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
118	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
119	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
120	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
121	ATCC # 98923	SignalTag 44-66
122	ATCC # 98920	SignalTag 67-90
123	ATCC # 98920	SignalTag 67-90
124	ATCC # 98922	SignalTag 91-94, 96, 97, 99-107, 109-112 et 114-120
125	ECACC # 98121506	SignalTag 11121998
126	ECACC # 98121506	SignalTag 11121998
127	ECACC # 98121506	SignalTag 11121998
128	ECACC # 98121506	SignalTag 11121998
129	ECACC # 98121506	SignalTag 11121998
130	ECACC # 98121506	SignalTag 11121998

CONT. TABLE VI

131	ECACC # 98121506	SignalTag 11121998
132	ECACC # 98121506	SignalTag 11121998
133	ECACC # 98121506	SignalTag 11121998
134	ECACC # 98121506	SignalTag 11121998
135	ECACC # 98121506	SignalTag 11121998
136	ECACC # 98121506	SignalTag 11121998
137	ECACC # 98121506	SignalTag 11121998
138	ECACC # 98121506	SignalTag 11121998
139	ECACC # 98121506	SignalTag 11121998
140	ECACC # 98121506	SignalTag 11121998

TABLE VII

INTERNAL DESIGNATION NUMBER	SEQ ID NO.	TYPE OF SEQUENCE
20-5-2-C3-CL0_4	40	DNA
20-8-4-A11-CL2_6	41	DNA
21-1-4-F2-CL11_1	42	DNA
22-11-2-H9-CL1_1	43	DNA
25-7-3-D4-CL0_2	44	DNA
26-27-3-D7-CL0_1	45	DNA
26-35-4-H9-CL1_1	46	DNA
26-45-2-C4-CL2_6	47	DNA
27-1-2-B3-CL0_1	48	DNA
27-1-2-B3-CL0_2	49	DNA
27-19-3-G7-CL11_2	50	DNA
33-10-4-E2-CL13_4	51	DNA
33-10-4-H2-CL2_2	52	DNA
33-110-4-A5-CL1_1	53	DNA
33-13-1-C1-CL1_1	54	DNA
33-30-2-A6-CL0_1	55	DNA
33-35-4-F4-CL1_2	56	DNA
33-35-4-G1-CL1_2	57	DNA
33-36-3-E2-CL1_1	58	DNA
33-36-3-E2-CL1_2	59	DNA
33-36-3-F2-CL2_2	60	DNA
33-4-2-G5-CL2_1	61	DNA
33-49-1-H4-CL1_1	62	DNA
33-66-2-B10-CL4_1	63	DNA
33-97-4-G8-CL2_2	64	DNA
33-98-4-C1-CL1_3	65	DNA
47-14-1-C3-CL0_5	66	DNA
47-15-1-E11-CL0_1	67	DNA
47-15-1-H8-CL0_2	68	DNA

CONT. TABLE VII

48-1-1-H7-CL0_1	69	DNA
48-1-1-H7-CL0_4	70	DNA
48-1-1-H7-CL0_5	71	DNA
48-3-1-H9-CL0_6	72	DNA
48-54-1-G9-CL2_1	73	DNA
48-54-1-G9-CL3_1	74	DNA
48-7-4-H2-CL2_2	75	DNA
51-11-3-D5-CL1_3	76	DNA
51-11-3-G9-CL0_1	77	DNA
51-15-4-A12-CL11_3	78	DNA
51-17-4-A4-CL3_1	79	DNA
51-2-3-F10-CL1_5	80	DNA
51-2-4-F5-CL11_2	81	DNA
51-27-4-F2-CL0_2	82	DNA
51-34-3-F8-CL0_2	83	DNA
57-1-4-E2-CL1_2	84	DNA
57-19-2-G8-CL2_1	85	DNA
57-27-3-G10-CL2_2	86	DNA
58-33-3-B4-CL1_2	87	DNA
58-34-3-C9-CL1_2	88	DNA
58-4-4-G2-CL2_1	89	DNA
58-48-1-G3-CL2_4	90	DNA
58-6-1-H4-CL1_1	91	DNA
60-12-1-E11-CL1_2	92	DNA
65-4-4-H3-CL1_1	93	DNA
74-5-1-E4-CL1_2	94	DNA
76-13-3-A9-CL1_2	95	DNA
76-16-1-D6-CL1_1	96	DNA
76-28-3-A12-CL1_5	97	DNA
76-42-2-F3-CL0_1	98	DNA
77-16-4-G3-CL1_3	99	DNA

CONT. TABLE VII

77-39-4-H4-CL11_4	100	DNA
78-24-3-H4-CL2_1	101	DNA
78-27-3-D1-CL1_6	102	DNA
78-28-3-D2-CL0_2	103	DNA
78-7-1-G5-CL2_6	104	DNA
84-3-1-G10-CL11_6	105	DNA
58-48-4-E2-CL0_1	106	DNA
23-12-2-G6-CL1_2	107	DNA
25-8-4-B12-CL0_5	108	DNA
26-44-3-C5-CL2_1	109	DNA
27-1-2-B3-CL0_3	110	DNA
30-12-3-G5-CL0_1	111	DNA
33-106-2-F10-CL1_3	112	DNA
33-28-4-D1-CL0_1	113	DNA
33-31-3-C8-CL2_1	114	DNA
48-24-1-D2-CL3_2	115	DNA
48-46-4-A11-CL1_4	116	DNA
51-1-4-C1-CL0_2	117	DNA
51-39-3-H2-CL1_2	118	DNA
51-42-3-F9-CL1_1	119	DNA
51-5-3-G2-CL0_4	120	DNA
57-18-4-H5-CL2_1	121	DNA
76-23-3-G8-CL1_1	122	DNA
76-23-3-G8-CL1_3	123	DNA
78-8-3-E6-CL0_1	124	DNA
19-10-1-C2-CL1_3	125	DNA
33-11-1-B11-CL1_2	126	DNA
33-113-2-B8-CL1_2	127	DNA
33-19-1-C11-CL1_1	128	DNA
33-61-2-F6-CL0_2	129	DNA
47-4-4-C6-CL2_2	130	DNA

CONT. TABLE VII

48-54-1-G9-CL1_1	131	DNA
51-43-3-G3-CL0_1	132	DNA
55-1-3-D11-CL0_1	133	DNA
58-14-2-D3-CL1_2	134	DNA
58-35-2-B6-CL2_3	135	DNA
76-18-1-F6-CL1_1	136	DNA
76-23-3-G8-CL2_2	137	DNA
76-30-3-B7-CL1_1	138	DNA
78-21-3-G7-CL2_1	139	DNA
58-45-4-B11-CL13_2	140	DNA
20-6-1-D11-FL2	242	DNA
20-8-4-A11-FL2	243	DNA
22-6-2-C1-FL2	244	DNA
22-11-2-H9-FL1	245	DNA
23-8-3-B1-FL1	246	DNA
24-3-3-C6-FL1	247	DNA
24-4-1-H3-FL1	248	DNA
26-45-2-C4-FL2	249	DNA
26-48-1-H10-FL1	250	DNA
26-49-1-A5-FL2	251	DNA
30-6-4-E3-FL3	252	DNA
33-6-1-G11-FL1	253	DNA
33-8-1-A3-FL2	254	DNA
33-11-3-C6-FL1	255	DNA
33-14-4-E1-FL1	256	DNA
33-21-2-D5-FL1	257	DNA
33-26-4-E10-FL1	258	DNA
33-27-1-E11-FL1	259	DNA
33-28-4-D1-FL1	260	DNA
33-28-4-E2-FL2	261	DNA
33-30-4-C4-FL1	262	DNA

CONT. TABLE VII

33-35-4-F4-FL1	263	DNA
33-36-3-F2-FL2	264	DNA
33-52-4-F9-FL2	265	DNA
33-52-4-H3-FL1	266	DNA
33-59-1-B7-FL1	267	DNA
33-71-1-A8-FL1	268	DNA
33-72-2-B2-FL1	269	DNA
33-105-2-C3-FL1	270	DNA
33-107-4-C3-FL1	271	DNA
33-110-2-G4-FL1	272	DNA
47-7-4-D2-FL2	273	DNA
47-10-2-G12-FL1	274	DNA
47-14-3-D8-FL1	275	DNA
47-18-3-C2-FL1	276	DNA
47-18-3-G5-FL2	277	DNA
47-18-4-E3-FL2	278	DNA
48-3-1-H9-FL3	279	DNA
48-4-2-H3-FL1	280	DNA
48-6-1-C9-FL1	281	DNA
48-7-4-H2-FL2	282	DNA
48-8-1-D8-FL3	283	DNA
48-13-3-H8-FL1	284	DNA
48-19-3-A7-FL1	285	DNA
48-19-3-G1-FL1	286	DNA
48-25-4-D8-FL1	287	DNA
48-21-4-H4-FL1	288	DNA
48-26-3-B8-FL2	289	DNA
48-29-1-E2-FL1	290	DNA
48-31-3-F7-FL1	291	DNA
48-47-3-A5-FL1	292	DNA
51-1-1-G12-FL1	293	DNA



CONT. TABLE VII

51-1-4-E9-FL3	294	DNA
51-1-4-E9-FL2	295	DNA
51-2-1-E10-FL1	296	DNA
51-2-3-F10-FL1	297	DNA
51-2-4-F5-FL1	298	DNA
51-3-3-B10-FL2	299	DNA
51-3-3-B10-FL3	300	DNA
51-7-3-G3-FL1	301	DNA
51-10-3-D11-FL1	302	DNA
51-11-3-D5-FL1	303	DNA
51-13-1-F7-FL3	304	DNA
51-15-4-H10-FL1	305	DNA
51-17-4-A4-FL1	306	DNA
51-18-1-C3-FL1	307	DNA
51-25-3-F3-FL1	308	DNA
51-27-1-E8-FL1	309	DNA
51-28-2-G1-FL2	310	DNA
51-39-3-H2-FL1	311	DNA
51-42-3-F9-FL1	312	DNA
51-44-4-H4-FL1	313	DNA
55-1-3-H10-FL1	314	DNA
55-5-4-A6-FL1	315	DNA
58-26-3-D1-FL1	316	DNA
57-18-1-D5-FL1	317	DNA
57-27-3-A11-FL1	318	DNA
57-27-3-G10-FL2	319	DNA
58-10-3-D12-FL1	320	DNA
58-11-1-G10-FL1	321	DNA
58-11-2-G8-FL2	322	DNA
58-36-3-A9-FL2	323	DNA
58-38-1-A2-FL2	324	DNA

CONT. TABLE VII

58-38-1-E5-FL1	325	DNA
58-44-2-B3-FL3	326	DNA
58-45-3-H11-FL1	327	DNA
58-53-2-B12-FL2	328	DNA
59-9-4-A10-FL1	329	DNA
60-16-3-A6-FL1	330	DNA
60-17-3-G8-FL2	331	DNA
62-5-4-B10-FL1	332	DNA
65-4-4-H3-FL1	333	DNA
74-3-1-B9-FL1	334	DNA
76-4-1-G5-FL1	335	DNA
76-7-3-A12-FL1	336	DNA
76-16-4-C9-FL3	337	DNA
76-30-3-B7-FL1	338	DNA
77-5-1-C2-FL1	339	DNA
77-5-4-E7-FL1	340	DNA
77-11-1-A3-FL1	341	DNA
77-16-3-D7-FL1	342	DNA
77-16-4-G3-FL1	343	DNA
77-25-1-A6-FL1	344	DNA
77-26-2-F2-FL3	345	DNA
78-6-2-E3-FL2	346	DNA
78-7-1-G5-FL2	347	DNA
78-16-2-C2-FL1	348	DNA
78-18-3-B4-FL3	349	DNA
78-20-1-G11-FL1	350	DNA
78-22-3-E10-FL1	351	DNA
78-24-2-B8-FL1	352	DNA
78-24-3-A8-FL1	353	DNA
78-24-3-H4-FL2	354	DNA
78-25-1-F11-FL1	355	DNA

CONT. TABLE VII

78-26-1-B5-FL1	356	DNA
78-27-3-D1-FL1	357	DNA
78-29-1-B2-FL1	358	DNA
78-29-4-B6-FL1	359	DNA
14-1-3-E6-FL1	360	DNA
30-9-1-G8-FL2	361	DNA
33-10-4-H2-FL2	362	DNA
33-10-4-H2-FL1	363	DNA
74-10-3-C9-FL2	364	DNA
33-97-4-G8-FL3	365	DNA
33-97-4-G8-FL2	366	DNA
33-104-4-H4-FL1	367	DNA
47-2-3-B3-FL1	368	DNA
47-37-4-G11-FL1	369	DNA
57-25-1-F10-FL2	370	DNA
58-19-3-D3-FL1	371	DNA
58-34-3-C9-FL2	372	DNA
58-48-4-E2-FL2	373	DNA
76-21-1-C4-FL1	374	DNA
78-26-2-H7-FL1	375	DNA
77-20-2-E11-FL1	376	DNA
47-1-3-F7-FL2	377	DNA
20-5-2-C3-CL0_4	141	PRT
20-8-4-A11-CL2_6	142	PRT
21-1-4-F2-CL11_1	143	PRT
22-11-2-H9-CL1_1	144	PRT
25-7-3-D4-CL0_2	145	PRT
26-27-3-D7-CL0_1	146	PRT
26-35-4-H9-CL1_1	147	PRT
26-45-2-C4-CL2_6	148	PRT
27-1-2-B3-CL0_1	149	PRT

CONT. TABLE VII

27-1-2-B3-CL0_2	150	PRT
27-19-3-G7-CL11_2	151	PRT
33-10-4-E2-CL13_4	152	PRT
33-10-4-H2-CL2_2	153	PRT
33-110-4-A5-CL1_1	154	PRT
33-13-1-C1-CL1_1	155	PRT
33-30-2-A6-CL0_1	156	PRT
33-35-4-F4-CL1_2	157	PRT
33-35-4-G1-CL1_2	158	PRT
33-36-3-E2-CL1_1	159	PRT
33-36-3-E2-CL1_2	160	PRT
33-36-3-F2-CL2_2	161	PRT
33-4-2-G5-CL2_1	162	PRT
33-49-1-H4-CL1_1	163	PRT
33-66-2-B10-CL4_1	164	PRT
33-97-4-G8-CL2_2	165	PRT
33-98-4-C1-CL1_3	166	PRT
47-14-1-C3-CL0_5	167	PRT
47-15-1-E11-CL0_1	168	PRT
47-15-1-H8-CL0_2	169	PRT
48-1-1-H7-CL0_1	170	PRT
48-1-1-H7-CL0_4	171	PRT
48-1-1-H7-CL0_5	172	PRT
48-3-1-H9-CL0_6	173	PRT
48-54-1-G9-CL2_1	174	PRT
48-54-1-G9-CL3_1	175	PRT
48-7-4-H2-CL2_2	176	PRT
51-11-3-D5-CL1_3	177	PRT
51-11-3-G9-CL0_1	178	PRT
51-15-4-A12-CL11_3	179	PRT
51-17-4-A4-CL3_1	180	PRT

CONT. TABLE VII

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51-2-4-F5-CL11_2	182	PRT
51-27-4-F2-CL0_2	183	PRT
51-34-3-F8-CL0_2	184	PRT
57-1-4-E2-CL1_2	185	PRT
57-19-2-G8-CL2_1	186	PRT
57-27-3-G10-CL2_2	187	PRT
58-33-3-B4-CL1_2	188	PRT
58-34-3-C9-CL1_2	189	PRT
58-4-4-G2-CL2_1	190	PRT
58-48-1-G3-CL2_4	191	PRT
58-6-1-H4-CL1_1	192	PRT
60-12-1-E11-CL1_2	193	PRT
65-4-4-H3-CL1_1	194	PRT
74-5-1-E4-CL1_2	195	PRT
76-13-3-A9-CL1_2	196	PRT
76-16-1-D6-CL1_1	197	PRT
76-28-3-A12-CL1_5	198	PRT
76-42-2-F3-CL0_1	199	PRT
77-16-4-G3-CL1_3	200	PRT
77-39-4-H4-CL11_4	201	PRT
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78-27-3-D1-CL1_6	203	PRT
78-28-3-D2-CL0_2	204	PRT
78-7-1-G5-CL2_6	205	PRT
84-3-1-G10-CL11_6	206	PRT
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26-44-3-C5-CL2_1	210	PRT
27-1-2-B3-CL0_3	211	PRT

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33-31-3-C8-CL2_1	215	PRT
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48-46-4-A11-CL1_4	217	PRT
51-1-4-C1-CL0_2	218	PRT
51-39-3-H2-CL1_2	219	PRT
51-42-3-F9-CL1_1	220	PRT
51-5-3-G2-CL0_4	221	PRT
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76-23-3-G8-CL1_3	224	PRT
78-8-3-E6-CL0_1	225	PRT
19-10-1-C2-CL1_3	226	PRT
33-11-1-B11-CL1_2	227	PRT
33-113-2-B8-CL1_2	228	PRT
33-19-1-C11-CL1_1	229	PRT
33-61-2-F6-CL0_2	230	PRT
47-4-4-C6-CL2_2	231	PRT
48-54-1-G9-CL1_1	232	PRT
51-43-3-G3-CL0_1	233	PRT
55-1-3-D11-CL0_1	234	PRT
58-14-2-D3-CL1_2	235	PRT
58-35-2-B6-CL2_3	236	PRT
76-18-1-F6-CL1_1	237	PRT
76-23-3-G8-CL2_2	238	PRT
76-30-3-B7-CL1_1	239	PRT
78-21-3-G7-CL2_1	240	PRT
58-45-4-B11-CL13_2	241	PRT
20-6-1-D11-FL2	378	PRT

CONT. TABLE VII

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24-4-1-H3-FL1	384	PRT
26-45-2-C4-FL2	385	PRT
26-48-1-H10-FL1	386	PRT
26-49-1-A5-FL2	387	PRT
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33-8-1-A3-FL2	390	PRT
33-11-3-C6-FL1	391	PRT
33-14-4-E1-FL1	392	PRT
33-21-2-D5-FL1	393	PRT
33-26-4-E10-FL1	394	PRT
33-27-1-E11-FL1	395	PRT
33-28-4-D1-FL1	396	PRT
33-28-4-E2-FL2	397	PRT
33-30-4-C4-FL1	398	PRT
33-35-4-F4-FL1	399	PRT
33-36-3-F2-FL2	400	PRT
33-52-4-F9-FL2	401	PRT
33-52-4-H3-FL1	402	PRT
33-59-1-B7-FL1	403	PRT
33-71-1-A8-FL1	404	PRT
33-72-2-B2-FL1	405	PRT
33-105-2-C3-FL1	406	PRT
33-107-4-C3-FL1	407	PRT
33-110-2-G4-FL1	408	PRT
47-7-4-D2-FL2	409	PRT

CONT. TABLE VII

47-10-2-G12-FL1	410	PRT
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47-18-3-C2-FL1	412	PRT
47-18-3-G5-FL2	413	PRT
47-18-4-E3-FL2	414	PRT
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48-4-2-H3-FL1	416	PRT
48-6-1-C9-FL1	417	PRT
48-7-4-H2-FL2	418	PRT
48-8-1-D8-FL3	419	PRT
48-13-3-H8-FL1	420	PRT
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48-25-4-D8-FL1	423	PRT
48-21-4-H4-FL1	424	PRT
48-26-3-B8-FL2	425	PRT
48-29-1-E2-FL1	426	PRT
48-31-3-F7-FL1	427	PRT
48-47-3-A5-FL1	428	PRT
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51-1-4-E9-FL3	430	PRT
51-1-4-E9-FL2	431	PRT
51-2-1-E10-FL1	432	PRT
51-2-3-F10-FL1	433	PRT
51-2-4-F5-FL1	434	PRT
51-3-3-B10-FL2	435	PRT
51-3-3-B10-FL3	436	PRT
51-7-3-G3-FL1	437	PRT
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51-11-3-D5-FL1	439	PRT
51-13-1-F7-FL3	440	PRT



CONT. TABLE VII

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51-28-2-G1-FL2	446	PRT
51-39-3-H2-FL1	447	PRT
51-42-3-F9-FL1	448	PRT
51-44-4-H4-FL1	449	PRT
55-1-3-H10-FL1	450	PRT
55-5-4-A6-FL1	451	PRT
58-26-3-D1-FL1	452	PRT
57-18-1-D5-FL1	453	PRT
57-27-3-A11-FL1	454	PRT
57-27-3-G10-FL2	455	PRT
58-10-3-D12-FL1	456	PRT
58-11-1-G10-FL1	457	PRT
58-11-2-G8-FL2	458	PRT
58-36-3-A9-FL2	459	PRT
58-38-1-A2-FL2	460	PRT
58-38-1-E5-FL1	461	PRT
58-44-2-B3-FL3	462	PRT
58-45-3-H11-FL1	463	PRT
58-53-2-B12-FL2	464	PRT
59-9-4-A10-FL1	465	PRT
60-16-3-A6-FL1	466	PRT
60-17-3-G8-FL2	467	PRT
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65-4-4-H3-FL1	469	PRT
74-3-1-B9-FL1	470	PRT
76-4-1-G5-FL1	471	PRT

CONT. TABLE VII

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77-25-1-A6-FL1	480	PRT
77-26-2-F2-FL3	481	PRT
78-6-2-E3-FL2	482	PRT
78-7-1-G5-FL2	483	PRT
78-16-2-C2-FL1	484	PRT
78-18-3-B4-FL3	485	PRT
78-20-1-G11-FL1	486	PRT
78-22-3-E10-FL1	487	PRT
78-24-2-B8-FL1	488	PRT
78-24-3-A8-FL1	489	PRT
78-24-3-H4-FL2	490	PRT
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78-26-1-B5-FL1	492	PRT
78-27-3-D1-FL1	493	PRT
78-29-1-B2-FL1	494	PRT
78-29-4-B6-FL1	495	PRT
14-1-3-E6-FL1	496	PRT
30-9-1-G8-FL2	497	PRT
33-10-4-H2-FL2	498	PRT
33-10-4-H2-FL1	499	PRT
74-10-3-C9-FL2	500	PRT
33-97-4-G8-FL3	501	PRT
33-97-4-G8-FL2	502	PRT

CONT. TABLE VII

33-104-4-H4-FL1	503	PRT
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47-37-4-G11-FL1	505	PRT
57-25-1-F10-FL2	506	PRT
58-19-3-D3-FL1	507	PRT
58-34-3-C9-FL2	508	PRT
58-48-4-E2-FL2	509	PRT
76-21-1-C4-FL1	510	PRT
78-26-2-H7-FL1	511	PRT
77-20-2-E11-FL1	512	PRT
47-1-3-F7-FL2	513	PRT

**TABLE VIII**

ID	Locations	PROSITE Signature Name
195	110-121	Aldehyde dehydrogenases cysytine active site
221	28-37	ATP synthase alpha and beta subunits signature
223	171-181	Regulator of chromosome condensation (RCC1) signature 2
225	90-112	Phosphatidylethanolamine-binding protein family signature
226	10-34	Protein kinases ATP-binding region signature

WHAT IS CLAIMED IS:

1. A purified or isolated nucleic acid comprising the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary thereto.
2. A purified or isolated nucleic acid comprising at least 10 consecutive  
5 bases of the sequence of one of SEQ ID NOs: 40-140 and 242-377 or one of the sequences complementary thereto.
3. A purified or isolated nucleic acid comprising the full coding sequences of one of SEQ ID NOs: 40, 42-44, 46, 48, 49, 51, 53, 60, 62-72, 76-78, 80-83, 85-88, 90, 93, 94, 97, 99-102, 104, 107-125, 127, 132, 135-138, 140 and 242-377 wherein the  
10 full coding sequence comprises the sequence encoding signal peptide and the sequence encoding mature protein.
4. A purified or isolated nucleic acid comprising the nucleotides of one of SEQ ID NOs: 40-44, 46, 48, 49, 51-53, 55, 56, 58-72, 75-78, 80-88, 90, 93, 94, 97, 99-125, 127, 132, 133, 135-138, 140, and 242-377 which encode a mature protein.
- 15 5. A purified or isolated nucleic acid comprising the nucleotides of one of SEQ ID NOs: 40, 42-46, 48, 49, 51, 53, 57, 60, 62-73, 76-78, 80-83, 85-88, 90, 93-95, 97, 99-102, 104, 107-125, 127, 128, 130, 132, 134-140 and 242-377 which encode the signal peptide.
6. A purified or isolated nucleic acid encoding a polypeptide having the  
20 sequence of one of the sequences of SEQ ID NOs: 141-241 and 378-513.
7. A purified or isolated nucleic acid encoding a polypeptide having the sequence of a mature protein included in one of the sequences of SEQ ID NOs: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513.
- 25 8. A purified or isolated nucleic acid encoding a polypeptide having the sequence of a signal peptide included in one of the sequences of SEQ ID NOs: 141, 143-147, 149, 150, 152, 154, 158, 161, 163-174, 177-179, 181-184, 186-189, 191, 194-196, 198, 200-203, 205, 208-226, 228, 229, 231, 233, 235-241, and 378-513.

9. A purified or isolated protein comprising the sequence of one of SEQ ID NOs: 141-241 and 378-513.

10. A purified or isolated polypeptide comprising at least 10 consecutive amino acids of one of the sequences of SEQ ID NOs: 141-241 and 378-513.

5 11. An isolated or purified polypeptide comprising a signal peptide of one of the polypeptides of SEQ ID NOs: 141, 143-147, 149, 150, 152, 154, 158, 161, 163-174, 177-179, 181-184, 186-189, 191, 194-196, 198, 200-203, 205, 208-226, 228, 229, 231, 233, 235-241, and 378-513.

10 12. An isolated or purified polypeptide comprising a mature protein of one of the polypeptides of SEQ ID NOs: 141-145, 147, 149, 150, 152-154, 156, 157, 159-172, 176-179, 181-189, 191, 194, 195, 198, 200-226, 228, 233, 234, 236-239, 241 and 378-513.

13. A method of making a protein comprising one of the sequences of SEQ ID NO: 141-241 and 378-513, comprising the steps of:

15 obtaining a cDNA comprising one of the sequences of sequence of SEQ ID NO: 40-140 and 242-377;

inserting said cDNA in an expression vector such that said cDNA is operably linked to a promoter; and

20 introducing said expression vector into a host cell whereby said host cell produces the protein encoded by said cDNA.

14. The method of Claim 13, further comprising the step of isolating said protein.

15. A protein obtainable by the method of Claim 14.

16. A host cell containing a recombinant nucleic acid of Claim 1.

25 17. A purified or isolated antibody capable of specifically binding to a protein having the sequence of one of SEQ ID NOs: 141-241 and 378-513.

18. In an array of polynucleotides of at least 15 nucleotides in length, the improvement comprising inclusion in said array of at least one of the sequences of SEQ

ID NOs: 40-140 and 242-377, or one of the sequences complementary to the sequences of SEQ ID NOs: 40-140 and 242-377, or a fragment thereof of at least 15 consecutive nucleotides.

5 19. A purified or isolated nucleic acid of at least 15 bases capable of hybridizing under stringent conditions to the sequence of one of SEQ ID NOs: 40-140 and 242-377 or a sequence complementary to one of the sequences of SEQ ID NOs: 40-140 and 242-377.

10 20. A purified or isolated antibody capable of binding to a polypeptide comprising at least 10 consecutive amino acids of the sequence of one of SEQ ID NOs: 141-241 and 378-513.

## IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Applicant : Edwards, et al. )  
 )  
Appl. No. : Unknown )  
 )  
Filed : Herewith )  
 )  
For : EXTENDED cDNAs )  
 )  
Group Art Unit : Unknown )  
 )

---

SEQUENCE SUBMISSION

Assistant Commissioner for Patents  
Washington, D.C. 20231

Dear Sir:

A copy of the Sequence Listing in computer readable form as required by 37 C.F.R.  
§ 1.821(e) is submitted herewith.

As required by 37 C.F.R. § 1.821(f), the data on the enclosed disk is identical to the  
Sequence Listing in the application filed herewith.

Respectfully submitted,

KNOBBE, MARTENS, OLSON & BEAR, LLP

Dated: Dec. 17, 1998

By: Daniel Hart  
Daniel Hart  
Registration No. 40,637  
Attorney of Record  
620 Newport Center Drive  
Sixteenth Floor  
Newport Beach, CA 92660  
(619) 235-8550



# SEQUENCE LISTING

<110> Edwards, Jean-Baptiste Dumas Milne  
Duclert, Aymeric  
Bougueleret, Lydie

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aca gcc atc ttg gca gtg gct gtw ggt ttc cca gtc tct caa gac cag      161
Thr Ala Ile Leu Ala Val Ala Val Gly Phe Pro Val Ser Gln Asp Gln
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Xaa Phe Val Phe Pro Tyr Pro Tyr Pro Phe Arg Pro Leu Pro Pro Ile
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Pro Phe Pro Arg Phe Pro Trp Phe Arg Arg Xaa Phe Pro Ile Pro Ile
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<223> blastn  
  
<221> misc\_feature  
<222> 118..184  
<223> blastn  
  
<221> misc\_feature  
<222> 56..113  
<223> blastn  
  
<221> misc\_feature  
<222> 454..485  
<223> blastn  
  
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<222> 118..545  
<223> blastn

<221> misc\_feature  
<222> 65..369  
<223> blastn

<221> misc\_feature  
<222> 61..399  
<223> blastn

<221> misc\_feature  
<222> 408..458  
<223> blastn

<221> misc\_feature  
<222> 60..399  
<223> blastn

<221> misc\_feature

<222> 393..432

<223> blastn

<221> sig\_peptide

<222> 346..408

<223> Von Heijne matrix

<400> 19

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actccttttta gcataggggc ttcggcgcca gcggccagcg ctagtcggtc tggtaagtgc 60
ctgatgccga gttccgtctc tcgctctttt tcctgggtccc aggcaaagcg gasgnagatc 120
ctcaaacggc ctagtgcttc gcgcttcggg agaaaatcag cgggtctaatt aattcctctg 180
gtttgttgaa gcagttacca agaattctca accctttccc acaaaagcta attgagtaca 240
cgttcctgtt gagtacacgt tcctgttgat ttacaaaagg tgcaggtatg agcaggtctg 300
aagactaaca ttttgtgaag ttgtaaaaca gaaaacctgt tagaa atg tgg tgg ttt 357
```

Met Trp Trp Phe

-20

```
cag caa ggc ctc agt ttc ctt cct tca gcc ctt gta att tgg aca tct 405
Gln Gln Gly Leu Ser Phe Leu Pro Ser Ala Leu Val Ile Trp Thr Ser
```

-15

-10

-5

```
gct gct ttc ata ttt tca tac att act gca gta aca ctc cac cat ata 453
Ala Ala Phe Ile Phe Ser Tyr Ile Thr Ala Val Thr Leu His His Ile
```

1

5

10

15

```
gac ccg gct tta cct tat atc agt gac act ggt aca gta gct cca raa 501
Asp Pro Ala Leu Pro Tyr Ile Ser Asp Thr Gly Thr Val Ala Pro Xaa
```

20

25

30

```
aaa tgc tta ttt ggg gca atg cta aat att gcg gca gtt tta tgt caa 549
Lys Cys Leu Phe Gly Ala Met Leu Asn Ile Ala Ala Val Leu Cys Gln
```

35

40

45

```
aaa tagaaatcag gaarataatt caacttaaag aakttcattt catgaccaa 602
```

Lys

```
ctcttcaraa acatgtcttt acaagcatat ctcttgtatt gctttctaca ctgttgaatt 662
```

```
gtctggcaat atttctgcag tggaaaattt gatttarmta gttcttgact gataaatatg 722
```

```
gtaagggtggg cttttcccc tgtgtaattg gctactatgt cttactgagc caagttgtaw 782
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tttgaaataa aatgatatga gagtgcacaa aaaaaaaaaa 822
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<210> 20

<211> 21

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> 1..21

<223> Von Heijne matrix

score 5.5

seq SFLPSALVIWTSA/AF

<400> 20

Met Trp Trp Phe Gln Gln Gly Leu Ser Phe Leu Pro Ser Ala Leu Val

1

5

10

15

Ile Trp Thr Ser Ala

20

<210> 21  
 <211> 405  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> misc\_feature  
 <222> complement(103..398)  
 <223> blastn

<221> sig\_peptide  
 <222> 185..295  
 <223> Von Heijne matrix

<400> 21  
 atcaccttct tctccatcct tstctgggcc agtccccarc ccagtccttc tcctgacctg 60  
 ccagcccaaa gtcagccttc agcacgcgct tttctgcaca cagatattcc aggcctacct 120  
 ggcattccag gacctccgma atgatgctcc agtcccttac aagcgcttcc tggatgaggg 180  
 tggc atg gtg ctg acc acc ctc ccc ttg ccc tct gcc aac agc cct gtg 229  
 Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val  
 -35 -30 -25  
 aac atg ccc acc act ggc ccc aac agc ctg agt tat gct agc tct gcc 277  
 Asn Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala  
 -20 -15 -10  
 ctg tcc ccc tgt ctg acc gct cca aak tcc ccc cgg ctt gct atg atg 325  
 Leu Ser Pro Cys Leu Thr Ala Pro Xaa Ser Pro Arg Leu Ala Met Met  
 -5 1 5 10  
 cct gac aac taaatatacct tatccaaatc aataaarwra raatcctccc 374  
 Pro Asp Asn  
 tccaraaggg tttctaaaaa caaaaaaaaaa a 405

<210> 22  
 <211> 37  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> 1..37  
 <223> Von Heijne matrix  
 score 5.9  
 seq LSYASSALSPCLT/AP

<400> 22  
 Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val Asn  
 1 5 10 15  
 Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala Leu  
 20 25 30  
 Ser Pro Cys Leu Thr  
 35

<210> 23  
 <211> 496



<212> DNA  
<213> Homo sapiens

<220>  
<221> misc\_feature  
<222> 149..331  
<223> blastn

<221> misc\_feature  
<222> 328..485  
<223> blastn

<221> misc\_feature  
<222> complement(182..496)  
<223> blastn

<221> sig\_peptide  
<222> 196..240  
<223> Von Heijne matrix

<400> 23

aaaaaattgg tcccagtttt caccctgccg cagggctggc tggggagggc agcggtttag	60
attagccgtg gcctaggccg ttttaacgggg tgacacgagc ntgcagggcc gagtccaagg	120
cccggagata ggaccaaccg tcaggaatgc gaggaatggt tttcttcgga ctctatcgag	180
gcacacagac agacc atg ggg att ctg tct aca gtg aca gcc tta aca ttt	231
Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe	
-15 -10 -5	
gcc ara gcc ctg gac ggc tgc aga aat ggc att gcc cac cct gca agt	279
Ala Xaa Ala Leu Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser	
1 5 10	
gag aag cac aga ctc gag aaa tgt agg gaa ctc gag asc asc cac tcg	327
Glu Lys His Arg Leu Glu Lys Cys Arg Glu Leu Glu Xaa Xaa His Ser	
15 20 25	
gcc cca gga tca acc cas cac cga aga aaa aca acc aga aga aat tat	375
Ala Pro Gly Ser Thr Xaa His Arg Arg Lys Thr Thr Arg Arg Asn Tyr	
30 35 40 45	
act tca gcc tgaaatgaak ccgggatcaa atggttgctg atcaragccc	424
Ser Ser Ala	
atattttaaat tggaaaagtc aaattgasca ttattaaata aagcttggtt aatatgtctc	484
aaacaaaaaa aa	496

<210> 24  
<211> 15  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> 1..15  
<223> Von Heijne matrix  
score 5.5  
seq ILSTVTALTFAXA/LD

<400> 24  
Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe Ala Xaa Ala

1 5 10 15

<210> 25  
<211> 623  
<212> DNA  
<213> Homo sapiens

<220>  
<221> sig\_peptide  
<222> 49..96  
<223> Von Heijne matrix

<400> 25  
aaagatccct gcagcccggc aggagagaag gctgagcctt ctggcgtc atg gag agg 57  
Met Glu Arg  
-15  
ctc gtc cta acc ctg tgc acc ctc ccg ctg gct gtg gcg tct gct ggc 105  
Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala Ser Ala Gly  
-10 -5 1  
tgc gcc acg acg cca gct cgc aac ctg agc tgc tac cag tgc ttc aag 153  
Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln Cys Phe Lys  
5 10 15  
gtc agc agc tgg acg gag tgc ccg ccc acc tgg tgc agc ccg ctg gac 201  
Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser Pro Leu Asp  
20 25 30 35  
Caa gtc tgc atc tcc aac gag gtg gtc gtc tct ttt aaa tgg agt gta 249  
Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Lys Trp Ser Val  
40 45 50  
cgc gtc ctg ctc agc aaa cgc tgt gct ccc aga tgt ccc aac gac aac 297  
Arg Val Leu Leu Ser Lys Arg Cys Ala Pro Arg Cys Pro Asn Asp Asn  
55 60 65  
atg aak ttc gaa tgg tgc ccg gcc ccc atg gtg caa ggc gtg atc acc 345  
Met Xaa Phe Glu Trp Ser Pro Ala Pro Met Val Gln Gly Val Ile Thr  
70 75 80  
agg cgc tgc tgt tcc tgg gct ctc tgc aac agg gca ctg acc cca cag 393  
Arg Arg Cys Cys Ser Trp Ala Leu Cys Asn Arg Ala Leu Thr Pro Gln  
85 90 95  
gag ggg cgc tgg gcc ctg cra ggg ggg ctc ctg ctc cag gac cct tcg 441  
Glu Gly Arg Trp Ala Leu Xaa Gly Gly Leu Leu Leu Gln Asp Pro Ser  
100 105 110 115  
agg ggc ara aaa acc tgg gtg cgg cca cag ctg ggg ctc cca ctc tgc 489  
Arg Gly Xaa Lys Thr Trp Val Arg Pro Gln Leu Gly Leu Pro Leu Cys  
120 125 130  
ctt ccc awt tcc aac ccc ctc tgc cca rgg gaa acc cag gaa gga 534  
Leu Pro Xaa Ser Asn Pro Leu Cys Pro Xaa Glu Thr Gln Glu Gly  
135 140 145  
taacactgtg ggtgccccca cctgtgcatt gggaccacra cttcaccctc ttggaracaa 594  
taaactctca tgcccccaaa aaaaaaaaaa 623

<210> 26  
<211> 16  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> 1..16  
<223> Von Heijne matrix  
score 10.1  
seq LVLTLCTLPLAVA/SA

<400> 26  
Met Glu Arg Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala  
1 5 10 15

<210> 27  
<211> 848  
<212> DNA  
<213> Homo sapiens

<220>  
<221> sig\_peptide  
<222> 32..73  
<223> Von Heijne matrix

<400> 27  
aaactttgcct tgtgtttttcc accctgaaag a atg ttg tgg ctg ctc ttt ttt 52  
Met Leu Trp Leu Leu Phe Phe  
-10  
ctg gtg act gcc att cat gct gaa ctc tgt caa cca ggt gca gaa aat 100  
Leu Val Thr Ala Ile His Ala Glu Leu Cys Gln Pro Gly Ala Glu Asn  
-5 1 5  
gct ttt aaa gtg aga ctt agt atc aga aca gct ctg gga gat aaa gca 148  
Ala Phe Lys Val Arg Leu Ser Ile Arg Thr Ala Leu Gly Asp Lys Ala  
10 15 20 25  
tat gcc tgg gat acc aat gaa gaa tac ctc ttc aaa gcg atg gta gct 196  
Tyr Ala Trp Asp Thr Asn Glu Glu Tyr Leu Phe Lys Ala Met Val Ala  
30 35 40  
ttc tcc atg aga aaa gtt ccc aac aga gaa gca aca gaa att tcc cat 244  
Phe Ser Met Arg Lys Val Pro Asn Arg Glu Ala Thr Glu Ile Ser His  
45 50 55  
gtc cta ctt tgc aat gta acc cag agg gta tca ttc tgg ttt gtg gtt 292  
Val Leu Leu Cys Asn Val Thr Gln Arg Val Ser Phe Trp Phe Val Val  
60 65 70  
aca gac cct tca aaa aat cac acc ctt cct gct gtt gag gtg caa tca 340  
Thr Asp Pro Ser Lys Asn His Thr Leu Pro Ala Val Glu Val Gln Ser  
75 80 85  
gcc ata aga atg aac aag aac cgg atc aac aat gcc ttc ttt cta aat 388  
Ala Ile Arg Met Asn Lys Asn Arg Ile Asn Asn Ala Phe Phe Leu Asn  
90 95 100 105  
gac caa act ctg gaa ttt tta aaa atc cct tcc aca ctt gca cca ccc 436  
Asp Gln Thr Leu Glu Phe Leu Lys Ile Pro Ser Thr Leu Ala Pro Pro  
110 115 120  
atg gac cca tct gtg ccc atc tgg att att ata ttt ggt gtg ata ttt 484  
Met Asp Pro Ser Val Pro Ile Trp Ile Ile Ile Phe Gly Val Ile Phe  
125 130 135  
tgc atc atc ata gtt gca att gca cta ctg att tta tca ggg atc tgg 532  
Cys Ile Ile Ile Val Ala Ile Ala Leu Leu Ile Leu Ser Gly Ile Trp

```

      140      145      150
caa cgt ada ara aag aac aaa gaa cca tct gaa gtg gat gac gct gaa      580
Gln Arg Xaa Xaa Lys Asn Lys Glu Pro Ser Glu Val Asp Asp Ala Glu
      155      160      165
rat aak tgt gaa aac atg atc aca att gaa aat ggc atc ccc tct gat      628
Xaa Xaa Cys Glu Asn Met Ile Thr Ile Glu Asn Gly Ile Pro Ser Asp
      170      175      180      185
ccc ctg gac atg aag gga ggg cat att aat gat gcc ttc atg aca gag      676
Pro Leu Asp Met Lys Gly Gly His Ile Asn Asp Ala Phe Met Thr Glu
      190      195      200
gat gag agg ctc acc cct ctc tgaagggtg ttgttctgct tcctcaaraa      727
Asp Glu Arg Leu Thr Pro Leu
      205
attaaacatt tgtttctgtg tgactgctga gcatacctgaa ataccaagag cagatcatat      787
wttttgtttc accattcttc ttttgtaata aattttgaat gtgcttgaaa aaaaaaaaaa      847
c                                                                 848

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<210> 28
<211> 14
<212> PRT
<213> Homo sapiens

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<220>
<221> SIGNAL
<222> 1..14
<223> Von Heijne matrix
      score 10.7
      seq LWLLFFLVTAIHA/EL

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<400> 28
Met Leu Trp Leu Leu Phe Phe Leu Val Thr Ala Ile His Ala
      5      10

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<210> 29
<211> 25
<212> DNA
<213> Artificial Sequence

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<220>
<223> Oligonucleotide

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```

<400> 29
gggaagatgg agatagtatt gcctg      25

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```

<210> 30
<211> 26
<212> DNA
<213> Artificial Sequence

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```

<220>
<223> Oligonucleotide

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<400> 30  
ctgccatgta catgatagag agattc

26

<210> 31  
<211> 546  
<212> DNA  
<213> Homo sapiens

<220>  
<221> promoter  
<222> 1..517

<221> transcription start site  
<222> 518

<221> protein\_bind  
<222> 17..25  
<223> matinspector prediction  
name CMYB\_01  
score 0.983  
sequence tgtcagttg

<221> protein\_bind  
<222> complement(18..27)  
<223> matinspector prediction  
name MYOD\_Q6  
score 0.961  
sequence cccaactgac

<221> protein\_bind  
<222> complement(75..85)  
<223> matinspector prediction  
name S8\_01  
score 0.960  
sequence aatagaattag

<221> protein\_bind  
<222> 94..104  
<223> matinspector prediction  
name S8\_01  
score 0.966  
sequence aactaaattag

<221> protein\_bind  
<222> complement(129..139)  
<223> matinspector prediction  
name DELTAEF1\_01  
score 0.960  
sequence gcacacctcag

<221> protein\_bind  
<222> complement(155..165)  
<223> matinspector prediction  
name GATA\_C

score 0.964  
sequence agataaatcca

<221> protein\_bind  
<222> 170..178  
<223> matinspector prediction  
name CMYB\_01  
score 0.958  
sequence cttcagttg

<221> protein\_bind  
<222> 176..189  
<223> matinspector prediction  
name GATA1\_02  
score 0.959  
sequence ttgtagataggaca

<221> protein\_bind  
<222> 180..190  
<223> matinspector prediction  
name GATA\_C  
score 0.953  
sequence agataggacat

<221> protein\_bind  
<222> 284..299  
<223> matinspector prediction  
name TAL1ALPHA47\_01  
score 0.973  
sequence cataacagatggtaag

<221> protein\_bind  
<222> 284..299  
<223> matinspector prediction  
name TAL1BETA47\_01  
score 0.983  
sequence cataacagatggtaag

<221> protein\_bind  
<222> 284..299  
<223> matinspector prediction  
name TAL1BETA1TF2\_01  
score 0.978  
sequence cataacagatggtaag

<221> protein\_bind  
<222> complement (287..296)  
<223> matinspector prediction  
name MYOD\_Q6  
score 0.954  
sequence accatctgtt

<221> protein\_bind  
<222> complement (302..314)  
<223> matinspector prediction  
name GATA1\_04  
score 0.953

sequence tcaagataaagta

<221> protein\_bind  
<222> 393..405  
<223> matinspector prediction  
name IK1\_01  
score 0.963  
sequence agttgggaattcc

<221> protein\_bind  
<222> 393..404  
<223> matinspector prediction  
name IK2\_01  
score 0.985  
sequence agttgggaattc

<221> protein\_bind  
<222> 396..405  
<223> matinspector prediction  
name CREL\_01  
score 0.962  
sequence tgggaattcc

<221> protein\_bind  
<222> 423..436  
<223> matinspector prediction  
name GATA1\_02  
score 0.950  
sequence tcagtgatatggca

<221> protein\_bind  
<222> complement(478..489)  
<223> matinspector prediction  
name SRY\_02  
score 0.951  
sequence taaaacaaaaca

<221> protein\_bind  
<222> 486..493  
<223> matinspector prediction  
name E2F\_02  
score 0.957  
sequence tttagcgc

<221> protein\_bind  
<222> complement(514..521)  
<223> matinspector prediction  
name MZF1\_01  
score 0.975  
sequence tgagggga

<400> 31  
tgagtgcagt gttacatgtc agttgggtta agtttggtta tgcattcaa atcttctatg 60  
tcttgatttg cctgctaatt ctattatttc tggaactaaa ttagtttgat ggttctatta 120  
gttattgact gaggtgtgct aatctcccat tatgtggatt tatctatttc ttcagttgta 180  
gataggacat tgatagatac ataagtacca ggacaaaagc agggagatct tttttccaaa 240  
atcaggagaa aaaaatgaca tctggaaaac ctatagggaa aggcataaca gatggtaagg 300

atactttatc ttgagtagga gagccttcct gtggcaacgt ggagaagggg agaggtcgta	360
gaattgagga gtcagctcag ttagaagcag ggagttggga attccgttca tgtgatttag	420
catcagtgat atggcaaagt tgggactaag ggtagtgatc agaggggttaa aattgtgtgt	480
tttgttttag cgctgctggg gcatgcctt gggccccctc aaacagattc ccatgaatct	540
cttcat	546

<210> 32  
<211> 23  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Oligonucleotide

<400> 32  
gtaccagggg ctgtgaccat tgc 23

<210> 33  
<211> 24  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Oligonucleotide

<400> 33  
ctgtgaccat tgctcccaag agag 24

<210> 34  
<211> 861  
<212> DNA  
<213> Homo sapiens

<220>  
<221> promoter  
<222> 1..806

<221> transcription start site  
<222> 807

<221> protein\_bind  
<222> complement(60..70)  
<223> matinspector prediction  
name NFY\_Q6  
score 0.956  
sequence ggaccaatcat

<221> protein\_bind  
<222> 70..77  
<223> matinspector prediction  
name MZF1\_01



score 0.962  
sequence cctgggga

<221> protein\_bind  
<222> 124..132  
<223> matinspector prediction  
name CMYB\_01  
score 0.994  
sequence tgaccgttg

<221> protein\_bind  
<222> complement(126..134)  
<223> matinspector prediction  
name VMYB\_02  
score 0.985  
sequence tccaacggt

<221> protein\_bind  
<222> 135..143  
<223> matinspector prediction  
name STAT\_01  
score 0.968  
sequence ttcctggaa

<221> protein\_bind  
<222> complement(135..143)  
<223> matinspector prediction  
name STAT\_01  
score 0.951  
sequence ttccaggaa

<221> protein\_bind  
<222> complement(252..259)  
<223> matinspector prediction  
name MZF1\_01  
score 0.956  
sequence ttggggga

<221> protein\_bind  
<222> 357..368  
<223> matinspector prediction  
name IK2\_01  
score 0.965  
sequence gaatgggatttc

<221> protein\_bind  
<222> 384..391  
<223> matinspector prediction  
name MZF1\_01  
score 0.986  
sequence agagggga

<221> protein\_bind  
<222> complement(410..421)  
<223> matinspector prediction  
name SRY\_02  
score 0.955

sequence gaaaacaaaaca

<221> protein\_bind

<222> 592..599

<223> matinspector prediction

name MZF1\_01

score 0.960

sequence gaagggga

<221> protein\_bind

<222> 618..627

<223> matinspector prediction

name MYOD\_Q6

score 0.981

sequence agcatctgcc

<221> protein\_bind

<222> 632..642

<223> matinspector prediction

name DELTAEF1\_01

score 0.958

sequence tcccaccttcc

<221> protein\_bind

<222> complement(813..823)

<223> matinspector prediction

name S8\_01

score 0.992

sequence gaggcaattat

<221> protein\_bind

<222> complement(824..831)

<223> matinspector prediction

name MZF1\_01

score 0.986

sequence agagggga

<400> 34

tactataggg	cacgcgtggt	cgacggccgg	gctgttcttg	agcagagggc	atgtcagtaa	60
tgattggtcc	ctggggaagg	tctggctggc	tccagcacag	tgaggcattt	aggtatctct	120
cggtgaccgt	tggattcctg	gaagcagtag	ctgttctggt	tggatctggt	agggacaggg	180
ctcagagggc	taggcacgag	ggaaggtcag	aggagaaggs	aggsarggcc	cagtgagarg	240
ggagcatgcc	ttccccaac	cctggcttsc	ycttggyam	agggcgkty	tgggmacttr	300
aaytcagggc	ccaascagaa	scacaggccc	aktcntggct	smaagcacia	tagcctgaat	360
gggatttcag	gttagncagg	gtgagagggg	aggctctctg	gcttagtttt	gttttgtttt	420
ccaaatcaag	gtaacttget	cccttctgct	acgggccttg	gtcttggttt	gtcctcacc	480
agtcggaact	ccctaccact	ttcaggagag	tggtttttag	cccgtggggc	tgttctgttc	540
caagcagtg	gagaacatgg	ctggtagagg	ctctagctgt	gtgcggggcc	tgaaggggag	600
tgggttctcg	cccaaagagc	atctgccc	ttcccacctt	cccttctccc	accagaagct	660
tgcctgagct	gtttggacaa	aatccaaac	cccacttggc	tactctggcc	tggcttcagc	720
ttggaacca	atacctaggc	ttacaggcca	tcttgagcca	ggggcctctg	gaaattctct	780
tctgatggg	cctttagggt	tgggcacaaa	atataattgc	ctctcccctc	tcccattttc	840
tctcttggga	gcaatggtca	c				861

<210> 35

<211> 20  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Oligonucleotide

<400> 35  
ctgggatgga aggcacggtg

20

<210> 36  
<211> 20  
<212> DNA  
<213> Artificial Sequence

<220>  
<223> Oligonucleotide

<400> 36  
gagaccacac agctagacaa

20

<210> 37  
<211> 555  
<212> DNA  
<213> Homo sapiens

<220>  
<221> promoter  
<222> 1..500  
  
<221> transcription start site  
<222> 501

<221> protein\_bind  
<222> 191..206  
<223> matinspector prediction  
name ARNT\_01  
score 0.964  
sequence ggactcacgtgctgct

<221> protein\_bind  
<222> 193..204  
<223> matinspector prediction  
name NMYC\_01  
score 0.965  
sequence actcacgtgctg

<221> protein\_bind  
<222> 193..204  
<223> matinspector prediction  
name USF\_01  
score 0.985  
sequence actcacgtgctg

<221> protein\_bind  
<222> complement(193..204)  
<223> matinspector prediction  
name USF\_01  
score 0.985  
sequence cagcacgtgagt

<221> protein\_bind  
<222> complement(193..204)  
<223> matinspector prediction  
name NMYC\_01  
score 0.956  
sequence cagcacgtgagt

<221> protein\_bind  
<222> complement(193..204)  
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name MYCMAX\_02  
score 0.972  
sequence cagcacgtgagt

<221> protein\_bind  
<222> 195..202  
<223> matinspector prediction  
name USF\_C  
score 0.997  
sequence tcacgtgc

<221> protein\_bind  
<222> complement(195..202)  
<223> matinspector prediction  
name USF\_C  
score 0.991  
sequence gcacgtga

<221> protein\_bind  
<222> complement(210..217)  
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name MZF1\_01  
score 0.968  
sequence catgggga

<221> protein\_bind  
<222> 397..410  
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name ELK1\_02  
score 0.963  
sequence ctctccggaagcct

<221> protein\_bind  
<222> 400..409  
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name CETS1P54\_01  
score 0.974  
sequence tccggaagcc

<221> protein\_bind  
<222> complement(460..470)  
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name AP1\_Q4  
score 0.963  
sequence agtgactgaac

<221> protein\_bind  
<222> complement(460..470)  
<223> matinspector prediction  
name AP1FJ\_Q2  
score 0.961  
sequence agtgactgaac

<221> protein\_bind  
<222> 547..555  
<223> matinspector prediction  
name PADS\_C  
score 1.000  
sequence tgtggtctc

<400> 37  
ctatagggca cgcktggtcg acggcccggt ctggtctggt ctgtkgtgga gtcgggttga 60  
aggacagcat ttgtkacatc tgggtctactg caccttcct ctgccgtgca cttggccttt 120  
kawaagctca gcaccggtgc ccatcacagg gccggcagca cacacatccc attactcaga 180  
aggaaactgac ggactcacgt gctgctccgt ccccatgagc tcagtggacc tgtctatgta 240  
gagcagtcag acagtgcctg ggatagagtg agagttcagc cagtaaatcc aagtgattgt 300  
cattcctgtc tgcattagta actcccaacc tagatgtgaa aacttagttc tttctcatag 360  
gttgtctctgc ccatgggtccc actgcagacc caggcactct ccggaagcct ggaaatcacc 420  
cgtgtctctc gctgtctccc gctcacatcc cacacttgtg ttcagtcact gagttacaga 480  
ttttgctccc tcaatttctc ttgtcttagt cccatcctct gttcccttgg ccagtttgtc 540  
tagctgtgtg gtctc 555

<210> 38

<211> 19

<212> DNA

<213> Artificial Sequence

<220>

<223> Oligonucleotide

<400> 38

ggccatacac ttgagtgc

19

<210> 39

<211> 19

<212> DNA

<213> Artificial Sequence

<220>

<223> Oligonucleotide

<400> 39

atatagacaa acgcacacc

19

<210> 40

<211> 568

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 7..471

<221> sig\_peptide

<222> 7..99

<223> Von Heijne matrix

score 6.9

seq LLLVPSALSLLLA/LL

<221> polyA\_signal

<222> 537..542

<221> polyA\_site

<222> 554..568

<400> 40

gggacc atg ttc acc agc acc ggc tcc agt ggg ctc tac aag gcg cct	48
Met Phe Thr Ser Thr Gly Ser Ser Gly Leu Tyr Lys Ala Pro	
-30 -25 -20	
ctg tcg aag agc ctt ctg ctg gtc ccc agt gcc ctc tcc ctc ctg ctc	96
Leu Ser Lys Ser Leu Leu Leu Val Pro Ser Ala Leu Ser Leu Leu Leu	
-15 -10 -5	
gcc ctc ctc ctg cct cac tgc cag aag ccc ttt gtg tat gac ctt cac	144
Ala Leu Leu Leu Pro His Cys Gln Lys Pro Phe Val Tyr Asp Leu His	
1 5 10 15	
gca gtc aag aac gac ttc cag att tgg agg ttg ata tgt gga aga ata	192
Ala Val Lys Asn Asp Phe Gln Ile Trp Arg Leu Ile Cys Gly Arg Ile	
20 25 30	
att tgc ctt gat ttg aaa gat act ttc tgc agt agt ctg ctt att tat	240
Ile Cys Leu Asp Leu Lys Asp Thr Phe Cys Ser Ser Leu Leu Ile Tyr	
35 40 45	
aat ttt agg ata ttt gaa aga aga tat gga agc aga aaa ttt gca tcc	288
Asn Phe Arg Ile Phe Glu Arg Arg Tyr Gly Ser Arg Lys Phe Ala Ser	
50 55 60	
ttt ttg ctg ggt acc tgg gtt ttg tca gcc tta ttt gac ttt ctc ctc	336
Phe Leu Leu Gly Thr Trp Val Leu Ser Ala Leu Phe Asp Phe Leu Leu	
65 70 75	
att gaa gct atg cag tat ttc ttt ggc atc act gca gct agt aat ttg	384
Ile Glu Ala Met Gln Tyr Phe Phe Gly Ile Thr Ala Ala Ser Asn Leu	
80 85 90 95	
cct tct gga tta atc ttt tgt tgt gct ttt tgc tct gag act aaa ctc	432
Pro Ser Gly Leu Ile Phe Cys Cys Ala Phe Cys Ser Glu Thr Lys Leu	
100 105 110	
ttc tta tca aga caa gct atg gca gag aac ttt tcc atc taataaattt	481
Phe Leu Ser Arg Gln Ala Met Ala Glu Asn Phe Ser Ile	
115 120	
aagagtagat tcattctgtat gggttgagagt aggctctgac tatgtatatg tgtataataa	541

acctacatat ccaaaaaaaaa aaaaaaa

568

<210> 41  
<211> 569  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 168..332

<221> polyA\_signal  
<222> 557..562

<400> 41

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tgccgcgcct tcgcctgccg cggctgtcaa ctgcctccgg agcgcggcgc cgagcgcagg	120
gatacggcgc ccagcgggggt cagaaagcaa cattgaatgc agaagaa atg gcg gac	176
	Met Ala Asp
	1
ttc tac aag gaa ttt tta agt aaa aat ttt cag aag cgc atg tat tat	224
Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Gln Lys Arg Met Tyr Tyr	
5 10 15	
aac aga gat tgg tac aag cgc aat ttt gcc atc acc ttc ttc atg gga	272
Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile Thr Phe Phe Met Gly	
20 25 30 35	
aaa gtg gcc ctg gaa agg att tgg aac aag ctt aaa cag aaa caa aag	320
Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu Lys Gln Lys Gln Lys	
40 45 50	
aag agg agc aac taggagtcca ctctgaccca gccagagtcc aggtttccac	372
Lys Arg Ser Asn	
55	
aggaagcaga tggagctcct ttcacagggg ctctgagaaa aactggagcc gatctcaaga	432
agccccacat cttcctaagg ggccccatgg cctgtttggg ggcagggtag gtcctggggc	492
actgtggggc gcctgcctgc tgatgtgggc tctaggccag cttgttgtca cgtacgtggt	552
gtgaaataaa gcccaag	569

<210> 42  
<211> 895  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 51..251

<221> sig\_peptide  
<222> 51..110  
<223> Von Heijne matrix  
score 5.3  
seq ALIFGGFISLIGA/AF

<221> polyA\_signal

<222> 849..854

<221> polyA\_site

<222> 882..895

<400> 42

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ccgagagtgc cgggcggtcg gcgggtcagg gcagcccggg gcctgacgcc atg tcc      56
                                     Met Ser
                                     -20
cgg aac ctg cgc acc gcg ctc att ttc ggc ggc ttc atc tcc ctg atc      104
Arg Asn Leu Arg Thr Ala Leu Ile Phe Gly Gly Phe Ile Ser Leu Ile
      -15                    -10                    -5
ggc gcc gcc ttc tat ccc atc tac ttc cgg ccc cta atg aga ttg gag      152
Gly Ala Ala Phe Tyr Pro Ile Tyr Phe Arg Pro Leu Met Arg Leu Glu
      1                    5                    10
gag tac aag aag gaa caa gct ata aat cgg gct gga att gtt caa gag      200
Glu Tyr Lys Lys Glu Gln Ala Ile Asn Arg Ala Gly Ile Val Gln Glu
      15                    20                    25                    30
gat gtg cag cca cca ggg tta aaa gtg tgg tct gat cca ttt ggc agg      248
Asp Val Gln Pro Pro Gly Leu Lys Val Trp Ser Asp Pro Phe Gly Arg
      35                    40                    45
aaa tgagagggct gtcacagct ctgattaaga aaggagattt cttcatgctt      301
Lys
tcgattctgc atggggtaca gccagtcacc tcaccagaga atgacggctg gagaagaaaa      361
ctctgtaata ccataaataa gagtgttgtt aataaaagac tgtgcacaag gattaatatt      421
tcccttctta agtatcaaaa gaactctgga acaaattata ccattaggaa ggttttcatg      481
attcagttga ttttccaaaa atgaagctat ctcacccagc tgggtttgga ggagcaatct      541
gcttattatt ctgtcgttac cacttactca agcgagctgt gatatgaata caagcaacca      601
gtgggctcgg gaagggtccgg gtctcttctg ccatcttcca gataagagat ttcagtaaaa      661
aactgccatg ctgagctgcc ttatagagct cttcgaaaat gttcgagttg ataaagctct      721
ttgaggacaa ggtacttcgt gcacctcatg ctgaagattg caccatgttg gaagataaat      781
atgaagcaag tcaaactaga tgcatacact tgtgtagaaa tcaataatca attaatagaa      841
gtgaaaaaat agacattaag atgattttatt tccactttgc aaaaaaaaaa aaaa      895
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<210> 43

<211> 691

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 20..613

<221> sig\_peptide

<222> 20..82

<223> Von Heijne matrix

score 10

seq LWALAMVTRPASA/AP

<400> 43

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ataccttaga ccctcagtc atg cca gtg cct gct ctg tgc ctg ctc tgg gcc      52
                                     Met Pro Val Pro Ala Leu Cys Leu Leu Trp Ala
                                     -20                    -15
ctg gca atg gtg acc cgg cct gcc tca gcg gcc ccc atg ggc ggc cca      100
Leu Ala Met Val Thr Arg Pro Ala Ser Ala Ala Pro Met Gly Gly Pro
```



```

-10          -5          1          5
gaa ctg gca cag cat gag gag ctg acc ctg ctc ttc cat ggg acc ctg      148
Glu Leu Ala Gln His Glu Glu Leu Thr Leu Leu Phe His Gly Thr Leu
          10          15          20
cag ctg ggc cag gcc ctc aac ggt gtg tac agg acc acg gag gga tgg      196
Gln Leu Gly Gln Ala Leu Asn Gly Val Tyr Arg Thr Thr Glu Gly Trp
          25          30          35
ctg aca aag gcc agg aac agc ctg ggt ctc tat ggc cgc aca ata gaa      244
Leu Thr Lys Ala Arg Asn Ser Leu Gly Leu Tyr Gly Arg Thr Ile Glu
          40          45          50
ctc ctg ggg cag gag gtc agc cgg ggc cgg gat gca gcc cag gaa ctt      292
Leu Leu Gly Gln Glu Val Ser Arg Gly Arg Asp Ala Ala Gln Glu Leu
          55          60          65          70
cgg gca agc ctg ttg gag act cag atg gag gag gat att ctg cag ctg      340
Arg Ala Ser Leu Leu Glu Thr Gln Met Glu Glu Asp Ile Leu Gln Leu
          75          80          85
cag gca gag gcc aca gct gag gtg ctg ggg gag gtg gcc cag gca cag      388
Gln Ala Glu Ala Thr Ala Glu Val Leu Gly Glu Val Ala Gln Ala Gln
          90          95          100
aag gtg cta cgg gac agc gtg cag cgg cta gaa gtc cag ctg agg agc      436
Lys Val Leu Arg Asp Ser Val Gln Arg Leu Glu Val Gln Leu Arg Ser
          105          110          115
gcc tgg ctg ggc cct gcc tac cga gaa ttt gag gtc tta aag gct cac      484
Ala Trp Leu Gly Pro Ala Tyr Arg Glu Phe Glu Val Leu Lys Ala His
          120          125          130
gct gac aag cag agc cac atc cta tgg gcc ctc aca ggc cac gtg cag      532
Ala Asp Lys Gln Ser His Ile Leu Trp Ala Leu Thr Gly His Val Gln
          135          140          145          150
cgg cag agg cgg gag atg gtg gca cag cag cat cgg ctg cga cag atc      580
Arg Gln Arg Arg Glu Met Val Ala Gln Gln His Arg Leu Arg Gln Ile
          155          160          165
cag gag aga ctc cac aca gcg gcg ctc cca gcc tgaatctgcc tggatggaac      633
Gln Glu Arg Leu His Thr Ala Ala Leu Pro Ala
          170          175
tgaggaccaa tcatgtctgca aggaacactt ccacgccccg tgaggcccct gtgcaggg      691

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<210> 44

<211> 458

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 12..416

<221> sig\_peptide

<222> 12..86

<223> Von Heijne matrix

score 4

seq LVVMVPLVGLIHL/GW

<221> polyA\_signal

<222> 425..430

<221> polyA\_site

<222> 445..458

<400> 44

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gctgaagtac t atg agc ctt cgg aac ttg tgg aga gac tac aaa gtt ttg      50
          Met Ser Leu Arg Asn Leu Trp Arg Asp Tyr Lys Val Leu
          -25          -20          -15
gtt gtt atg gtc cct tta gtt ggg ctc ata cat ttg ggg tgg tac aga      98
Val Val Met Val Pro Leu Val Gly Leu Ile His Leu Gly Trp Tyr Arg
          -10          -5          1
atc aaa agc agc cct gtt ttc caa ata cct aaa aac gac gac att cct      146
Ile Lys Ser Ser Pro Val Phe Gln Ile Pro Lys Asn Asp Asp Ile Pro
5          10          15          20
gag caa gat agt ctg gga ctt tca aat ctt cag aag agc caa atc cag      194
Glu Gln Asp Ser Leu Gly Leu Ser Asn Leu Gln Lys Ser Gln Ile Gln
          25          30          35
ggg aag nta gca ggc ttg caa tct tca ggt aaa gaa gca gct ttg aat      242
Gly Lys Xaa Ala Gly Leu Gln Ser Ser Gly Lys Glu Ala Ala Leu Asn
          40          45          50
ctg agc ttc ata tcg aaa gaa gag atg aaa aat acc agt tgg att aga      290
Leu Ser Phe Ile Ser Lys Glu Glu Met Lys Asn Thr Ser Trp Ile Arg
          55          60          65
aag aac tgg ctt ctt gta gct ggg ata tct ttc ata ggt gac cat ctt      338
Lys Asn Trp Leu Leu Val Ala Gly Ile Ser Phe Ile Gly Asp His Leu
          70          75          80
gga aca tac ttt ttg cag agg tct gca aag cag tct gta aaa ttt cag      386
Gly Thr Tyr Phe Leu Gln Arg Ser Ala Lys Gln Ser Val Lys Phe Gln
          85          90          95          100
tct caa agc aaa caa aag agt att gaa gag tgaagtaaaa taaatatttg      436
Ser Gln Ser Lys Gln Lys Ser Ile Glu Glu
          105          110
gaattactaa aaaaaaaaaa aa      458
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<210> 45

<211> 2036

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 276..1040

<221> sig\_peptide

<222> 276..485

<223> Von Heijne matrix

score 3.9

seq SVIGVMLAPFTAG/LS

<221> polyA\_site

<222> 2024..2036

<400> 45

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gatcctgggt gcagctcatc acaagcgtcg ggggtgcagca aaaccatcca ggctggacag      60
tggtctggaca gttccaagaa aagaaacgct tcaccgaaga agtcattgaa tacttccaga      120
agaaagtttag cccagtgcac ctgaaaatcc tgctgactag cgatgaagcc tggaagagat      180
tcgtgcgtgt ggctggattg cccaggggaag aagcagatgc tctctatgaa gctctgaaga      240
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atcttacacc atatgtggct attgaggaca aagac atg cag caa aaa gaa cag	293
Met Gln Gln Lys Glu Gln	
-70 -65	
cag ttt agg gag tgg ttt ttg aaa gag ttt cct caa atc aga tgg aag	341
Gln Phe Arg Glu Trp Phe Leu Lys Glu Phe Pro Gln Ile Arg Trp Lys	
-60 -55 -50	
att cag gag tcc ata gaa agg ctt cgt gtc att gca aat gag att gaa	389
Ile Gln Glu Ser Ile Glu Arg Leu Arg Val Ile Ala Asn Glu Ile Glu	
-45 -40 -35	
aag gtc cac aga ggc tgc gtc atc gcc aat gtg gtg tct ggc tcc act	437
Lys Val His Arg Gly Cys Val Ile Ala Asn Val Val Ser Gly Ser Thr	
-30 -25 -20	
ggc atc ctg tct gtc att ggc gtt atg ttg gca cca ttt aca gca ggg	485
Gly Ile Leu Ser Val Ile Gly Val Met Leu Ala Pro Phe Thr Ala Gly	
-15 -10 -5	
ctg agc ctg agc att act gca gct ggg gta ggg ctg gga ata gca tct	533
Leu Ser Leu Ser Ile Thr Ala Ala Gly Val Gly Leu Gly Ile Ala Ser	
1 5 10 15	
gcc acg gct ggg atc gcc tcc agc atc gtg gag aac aca tac aca agg	581
Ala Thr Ala Gly Ile Ala Ser Ser Ile Val Glu Asn Thr Tyr Thr Arg	
20 25 30	
tca gca gaa ctc aca gcc agc agg ctg act gca acc agc act gac caa	629
Ser Ala Glu Leu Thr Ala Ser Arg Leu Thr Ala Thr Ser Thr Asp Gln	
35 40 45	
attg gag gca tta agg gac att ctg cat gac atc aca ccc aat gtg ctt	677
Leu Glu Ala Leu Arg Asp Ile Leu His Asp Ile Thr Pro Asn Val Leu	
50 55 60	
acc ttt gca ctt gat ttt gac gaa gcc aca aaa atg att gcg aat gat	725
Ser Phe Ala Leu Asp Phe Asp Glu Ala Thr Lys Met Ile Ala Asn Asp	
65 70 75 80	
gtc cat aca ctc agg aga tct aaa gcc act gtt gga cgc cct ttg att	773
Val His Thr Leu Arg Arg Ser Lys Ala Thr Val Gly Arg Pro Leu Ile	
85 90 95	
gct tgg cga tat gta cct ata aat gtt gtt gag aca ctg aga aca cgt	821
Ala Trp Arg Tyr Val Pro Ile Asn Val Val Glu Thr Leu Arg Thr Arg	
100 105 110	
ggg gcc ccc acc cgg ata gtg aga aaa gta gcc cgg aac ctg ggc aag	869
Gly Ala Pro Thr Arg Ile Val Arg Lys Val Ala Arg Asn Leu Gly Lys	
115 120 125	
gcc act tca ggt gtc ctc gtt gtg ctg gat gta gtc aac ctt gtg caa	917
Ala Thr Ser Gly Val Leu Val Val Leu Asp Val Val Asn Leu Val Gln	
130 135 140	
gac tca ctg gac ttg cac aag ggg gaa aaa tcc gag tct gct gag ttg	965
Asp Ser Leu Asp Leu His Lys Gly Glu Lys Ser Glu Ser Ala Glu Leu	
145 150 155 160	
ctg agg cag tgg gct cag gag ctg gag gag aat ctc aat gag ctc acc	1013
Leu Arg Gln Trp Ala Gln Glu Leu Glu Glu Asn Leu Asn Glu Leu Thr	
165 170 175	
cat atc cat cag agt cta aaa gca ggc taggcccaat tgttgcgaggga	1060
His Ile His Gln Ser Leu Lys Ala Gly	
180 185	
agtcagggac cccaaacgga gggactggct gaagccatgg cagaagaacg tggattgtga	1120
agatttcatg gacatttatt agttcccaa attaatattt ttataatttc ctatgcctgt	1180
ctttaccgca atctctaaac acaaattgtg aagatttcat ggacacttat cacttcccca	1240
atcaataccc ttgtgatttc ttatgcctgt ctttacttta atctcctaata cctgtcagct	1300
gaggaggggtg tatgtcacct caggaccatg tgataattgc gttaactgca caaattgtag	1360
agcatgtgtg tttgaacaat atgaaatctg ggcaccttga aaaaagaaca ggataacagc	1420

aatcgttcag	gggataagag	agataaacctt	aaactctgac	caacagtgag	ccgggtggag	1480
cagagtcata	tttcttttct	ttcaaaagca	aatgggagaa	atatcgctga	attctttttc	1540
tcagcaagga	acatccctga	gaaagagaat	gcacccctga	gggtgggtct	ataaatggcc	1600
tccttgggtg	tggccatctt	ctatggtcga	gactgtaggg	atgaaataaa	ccccagtctc	1660
ccatagtgtc	cccaggctta	ttaggaagag	gaaattcccc	cctaataaat	tttggtcaga	1720
ccggttgctc	tcaaaaccct	gtctcctgat	aagatgttat	caatgacaat	ggtgcctgaa	1780
acctcattag	caattttaat	ttctccccgg	tctgtgggtc	ctgtgatctc	accctgcctc	1840
cacttgcctt	gtgatattct	attaccttgt	gaagtaggtg	atctttgtga	cccacaccct	1900
attcatacac	tccctcccct	tttggaagtc	cctaataaaa	acttgctggg	tttgcagctt	1960
gtgaggcatc	acggaaccta	ctgatgtgtg	atgtctcccc	tggacaccta	gctttaaaat	2020
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<210> 46

<211> 1276

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 443..619

<221> sig\_peptide

<222> 443..589

<223> Von Heijne matrix

score 7

seq LICVVCLYIVCRC/GS

<221> polyA\_site

<222> 1267..1276

<400> 46

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tgtaaataat	tggtagaaaa	attctactct	gctgtggaat	taccaagata	atatagacca	180
gaaactagct	gatcaaatta	atgatctcca	acaaactgta	atgtggctag	gggatcatat	240
agttagttta	gaatatagaa	tgcggttaca	atgtgattga	aatacctctg	atttttgcac	300
tactcctcat	ctgtgtaatg	aaacagagca	tgagtgggaa	aaagttaaga	gatattttaa	360
aggtcatact	agaaatttat	ctttggatat	tgcaaagcta	aaggaacaag	tatttcaagc	420
ccctcagata	catctgacac	ta atg cca gga act gaa gtg ctt gaa gga gct				472
		Met Pro Gly Thr Glu Val Leu Glu Gly Ala				
		-45			-40	
aca gac gga tta gca gct att aac ctg cta aaa tgg atc aag aca ctt						520
Thr Asp Gly Leu Ala Ala Ile Asn Leu Leu Lys Trp Ile Lys Thr Leu						
	-35		-30		-25	
gga ggc tct gtg att tca atg att gtg ctt tta atc tgt gtt gtt tgt						568
Gly Gly Ser Val Ile Ser Met Ile Val Leu Leu Ile Cys Val Val Cys						
	-20		-15		-10	
ctt tat ata gtc tgt aga tgc gga agc cac ctc tgg aga gaa agc cac						616
Leu Tyr Ile Val Cys Arg Cys Gly Ser His Leu Trp Arg Glu Ser His						
	-5	1	5			
cac tgagagcaag caatgatagc tgtggcgggtt ttgcaaaaag aaaagggaga						669
His						
10						
caagcgccca gctatagtta ccaataaagc atgggtactgg tattaaaata ggcatgtgtt						729
ctgttccaat ggaacagaat agagaacca gaaacaaagc caaatattta cagccaactg						789

atctctgaca	aagcaaacaa	aaacataaaag	tgggggaaagg	acaccctatt	ccacaaatag	849
tgcagggata	attggcaagc	cacatgtaga	aaaatgaagc	tggatcctcg	tctctcactt	909
tatacaaaaa	tcaactcaaa	atgggtcaaa	gtcttaactc	taagacctga	aaccataaca	969
attctagaaa	ataacattgg	aaaaactctt	ctagacattg	gtttaggcaa	aaagttcatg	1029
accaagaacc	caaaagcaaa	tgcaataaaa	aggaagataa	atagatggga	cctaattaag	1089
ctgaaaagct	tctgcatagc	aaaaggaata	atcagcagag	caaacagaca	accacagggg	1149
tggggagaaa	tatttgcaag	ctatgtatct	gacaatggac	taatatccag	aatctacaag	1209
gaattcaaac	aattagcaag	aaaaaacact	tgtatttgtt	ttgctctgta	aatcagcaaa	1269
aaaaaaa						1276

<210> 47

<211> 747

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 206..745

<400> 47

accagaagca	ggtgatttcc	gagctcagca	atgctcagct	cataatgatg	tcaagcacca	60
atggccagttt	tatgaatggc	ttcctgtgtc	taatgaccct	gacaacccat	gttcactcaa	120
gtgccaagcc	aaaggaacaa	ccctgggtgt	tgaactagca	cctaagggtct	tagatgggtac	180
gcgttgctat	acagaatctt	tggat	atg tgc atc agt	ggt tta tgc caa att		232
			Met Cys Ile Ser	Gly Leu Cys Gln Ile		
			1	5		
ggt ggc tgc gat cac cag ctg gga agc acc gtc aag gaa gat aac tgt						280
Val Gly Cys Asp His Gln Leu Gly Ser Thr Val Lys Glu Asp Asn Cys						
10	15	20	25			
ggg gtc tgc aac gga gat ggg tcc acc tgc cgg ctg gtc cga ggg cag						328
Gly Val Cys Asn Gly Asp Gly Ser Thr Cys Arg Leu Val Arg Gly Gln						
	30	35	40			
tat aaa tcc cag ctg tcc gca acc aaa tcg gat gat act gtg gtt gca						376
Tyr Lys Ser Gln Leu Ser Ala Thr Lys Ser Asp Asp Thr Val Val Ala						
	45	50	55			
att ccc tat gga agt aga cat att cgc ctt gtc tta aaa ggt cct gat						424
Ile Pro Tyr Gly Ser Arg His Ile Arg Leu Val Leu Lys Gly Pro Asp						
	60	65	70			
cac tta tat ctg gaa acc aaa acc ctg cag ggg act aaa ggt gaa aac						472
His Leu Tyr Leu Glu Thr Lys Thr Leu Gln Gly Thr Lys Gly Glu Asn						
	75	80	85			
agt ctg agc tcc aca gga act ttc ctt gtg gac aat tct agt gtg gac						520
Ser Leu Ser Ser Thr Gly Thr Phe Leu Val Asp Asn Ser Ser Val Asp						
	90	95	100	105		
ttc cag aaa ttt cca gac aaa gag ata ctg aga atg gct gga cca ctg						568
Phe Gln Lys Phe Pro Asp Lys Glu Ile Leu Arg Met Ala Gly Pro Leu						
	110	115	120			
aca gca gat ttc att gtc aag att cgt aac tcg ggc tcc gct gac agt						616
Thr Ala Asp Phe Ile Val Lys Ile Arg Asn Ser Gly Ser Ala Asp Ser						
	125	130	135			
aca gtc cag ttc atc ttc tat caa ccc atc atc cac cga tgg agg gag						664
Thr Val Gln Phe Ile Phe Tyr Gln Pro Ile Ile His Arg Trp Arg Glu						
	140	145	150			
acg gat ttc ttt cct tgc tca gca acc tgt gga gga ggt tat cag ctg						712
Thr Asp Phe Phe Pro Cys Ser Ala Thr Cys Gly Gly Gly Tyr Gln Leu						

155	160	165	
aca tcg gct gag tgc tac gat ctg agg agc aac cg			747
Thr Ser Ala Glu Cys Tyr Asp Leu Arg Ser Asn			
170	175	180	

<210> 48  
 <211> 561  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 36..521

<221> sig\_peptide  
 <222> 36..104  
 <223> Von Heijne matrix  
 score 7.4  
 seq VLLLAALPPVLLP/GA

<221> polyA\_signal  
 <222> 528..533  
 <221> polyA\_site  
 <222> 548..561

<400> 48	
gagcgctctt tcagcccgagg atcgcccgagg caggg atg ggc gac aag atc tgg	53
Met Gly Asp Lys Ile Trp	
-20	
ctg ccc ttc ccc gtg ctc ctt ctg gcc gct ctg cct ccg gtg ctg ctg	101
Leu Pro Phe Pro Val Leu Leu Leu Ala Ala Leu Pro Pro Val Leu Leu	
-15 -10 -5	
gcct ggg gcg gcc ggc ttc aca cct tcc ctc gat agc gac ttc acc ttt	149
Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu Asp Ser Asp Phe Thr Phe	
1 5 10 15	
acc ctt ccc gcc ggc cag aag gag tgc ttc tac cag ccc atg ccc ctg	197
Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe Tyr Gln Pro Met Pro Leu	
20 25 30	
aag gcc tcg ctg gag atc gag tac caa gtt tta gat gga gca gga tta	245
Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val Leu Asp Gly Ala Gly Leu	
35 40 45	
gat att gat ttc cat ctt gcc tct cca gaa ggc aaa acc tta gtt ttt	293
Asp Ile Asp Phe His Leu Ala Ser Pro Glu Gly Lys Thr Leu Val Phe	
50 55 60	
gaa caa aga aaa tca gat gga gtt cac act gta gag act gaa gtt ggt	341
Glu Gln Arg Lys Ser Asp Gly Val His Thr Val Glu Thr Glu Val Gly	
65 70 75	
gat tac atg ttc tgc ttt gac aat aca ttc agc acc att tct gag aag	389
Asp Tyr Met Phe Cys Phe Asp Asn Thr Phe Ser Thr Ile Ser Glu Lys	
80 85 90 95	
gtg att ttc ttt gaa tta atc ccg gat aat atg gga gaa cag gca caa	437
Val Ile Phe Phe Glu Leu Ile Pro Asp Asn Met Gly Glu Gln Ala Gln	
100 105 110	
gaa caa gaa gat tgg aag aaa tat att act ggc aca gat ata ttg gat	485

Glu Gln Glu Asp Trp Lys Lys Tyr Ile Thr Gly Thr Asp Ile Leu Asp	
115 120 125	
atg aaa ctg gaa gac atc ctg gtc agt atg gtc ttc taataaaata	531
Met Lys Leu Glu Asp Ile Leu Val Ser Met Val Phe	
130 135	
aaaattatta acagccaaaa aaaaaaaaaa	561

<210> 49  
 <211> 632  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 36..395

<221> sig\_peptide  
 <222> 36..104  
 <223> Von Heijne matrix  
 score 7.4  
 seq VLLLAALPPVLLP/GA

<221> polyA\_signal  
 <222> 599..604

<221> polyA\_site  
 <222> 619..632

<400> 49	
gacgcctctt tcagccccggg atcgccccag caggg atg ggc gac aag atc tgg	53
Met Gly Asp Lys Ile Trp	
-20	
ctg ccc ttc ccc gtg ctc ctt ctg gcc gct ctg cct ccg gtg ctg ctg	101
Leu Pro Phe Pro Val Leu Leu Leu Ala Ala Leu Pro Pro Val Leu Leu	
-15 -10 -5	
cct ggg gcg gcc ggc ttc aca cct tcc ctc gat agc gac ttc acc ttt	149
Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu Asp Ser Asp Phe Thr Phe	
1 5 10 15	
acc ctt ccc gcc ggc cag aag gag tgc ttc tac cag ccc atg ccc ctg	197
Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe Tyr Gln Pro Met Pro Leu	
20 25 30	
aag gcc tcg ctg gag atc gag tac caa gtt tta gat gga gca gga tta	245
Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val Leu Asp Gly Ala Gly Leu	
35 40 45	
gat att gat ttc cat ctt gcc tct cca gaa ggc aaa acc tta gtt ttt	293
Asp Ile Asp Phe His Leu Ala Ser Pro Glu Gly Lys Thr Leu Val Phe	
50 55 60	
gaa caa aga aaa tca gat gga gtt cac acg tgt ata aga agt aaa aat	341
Glu Gln Arg Lys Ser Asp Gly Val His Thr Cys Ile Arg Ser Lys Asn	
65 70 75	
ggg cca ggc act gcg gtt cac gcc tat aat ccc agc act ttc cga ggc	389
Gly Pro Gly Thr Ala Val His Ala Tyr Asn Pro Ser Thr Phe Arg Gly	
80 85 90 95	
caa gtg tagagactga agttggtgat tacatgttct gctttgacaa tacattcagc	445
Gln Val	

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accatttctg agaaggtgat tttctttgaa ttaatcctgg ataatatggg agaacaggca 505
caaggacaag aagattggaa gaaatatatt actggcacag atatattgga tatgaaactg 565
gaagacatcc tggtcagtat ggtcttctaa taaaataaaa attattaaca gccaaaaaaa 625
aaaaaaa 632
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<210> 50
<211> 370
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 21..41
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<221> polyA_signal
<222> 328..333
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<221> polyA_site
<222> 357..370
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<400> 50
ctgggacttc tggcctcaca atg gtt gag atg act ggg gtg tagcagtgcc 51
Met Val Glu Met Thr Gly Val
1 5
aagtcgaggc tgtgaaaggc cttccacctt tactctcgtg ctcgtgccct cccccattgt 111
taggagaagg gcatgctcag gccagcccat tagcccagga ggaggacaag aaacacacgg 171
agcagacaca agccacctca ccaaccacgc caaggctgtc ctgaattagc aaccctgaca 231
cgtgtgagca agtccaacgg acaccggaag atccacctag tcaagcccaa ccaagactgg 291
cagagctgcc aagctgacca cttaaggcgc atgaggaata aacactcgtt gctgcatgcc 351
attgcaaaaa aaaaaaaaaa 370
```

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<210> 51
<211> 994
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 35..631
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<221> sig_peptide
<222> 35..160
<223> Von Heijne matrix
score 8.6
seq ASLFLLLSLTVFS/IV
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<221> polyA_signal
<222> 901..906
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<221> polyA_site
<222> 979..994
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<400> 51
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ataattggag ctgcaaagca gatcgtgaca agag atg gac ggt cag aag aaa aat      55
                               Met Asp Gly Gln Lys Lys Asn
                               -40
tgg aag gac aag gtt gtt gac ctc ctg tac tgg aga gac att aag aag      103
Trp Lys Asp Lys Val Val Asp Leu Leu Tyr Trp Arg Asp Ile Lys Lys
-35          -30          -25          -20
act gga gtg gtg ttt ggt gcc agc cta ttc ctg ctg ctt tca ttg aca      151
Thr Gly Val Val Phe Gly Ala Ser Leu Phe Leu Leu Leu Ser Leu Thr
          -15          -10          -5
gta ttc agc att gtg agc gta aca gcc tac att gcc ttg gcc ctg ctc      199
Val Phe Ser Ile Val Ser Val Thr Ala Tyr Ile Ala Leu Ala Leu Leu
          1          5          10
tct gtg acc atc agc ttt agg ata tac aag ggt gtg atc caa gct atc      247
Ser Val Thr Ile Ser Phe Arg Ile Tyr Lys Gly Val Ile Gln Ala Ile
          15          20          25
cag aaa tca gat gaa ggc cac cca ttc agg gca tat ctg gaa tct gaa      295
Gln Lys Ser Asp Glu Gly His Pro Phe Arg Ala Tyr Leu Glu Ser Glu
          30          35          40          45
gtt gct ata tct gag gag ttg gtt cag aag tac agt aat tct gct ctt      343
Val Ala Ile Ser Glu Glu Leu Val Gln Lys Tyr Ser Asn Ser Ala Leu
          50          55          60
ggg cat gtg aac tgc acg ata aag gaa ctc agg cgc ctc ttc tta gtt      391
Gly His Val Asn Cys Thr Ile Lys Glu Leu Arg Arg Leu Phe Leu Val
          65          70          75
gat gat tta gtt gat tct ctg aag ttt gca gtg ttg atg tgg gta ttt      439
Asp Asp Leu Val Asp Ser Leu Lys Phe Ala Val Leu Met Trp Val Phe
          80          85          90
acc tat gtt ggt gcc ttg ttt aat ggt ctg aca cta ctg att ttg gct      487
Thr Tyr Val Gly Ala Leu Phe Asn Gly Leu Thr Leu Leu Ile Leu Ala
          95          100          105
ctc att tca ctc ttc agt gtt cct gtt att tat gaa cgg cat cag gca      535
Leu Ile Ser Leu Phe Ser Val Pro Val Ile Tyr Glu Arg His Gln Ala
          110          115          120          125
cag ata gat cat tat cta gta ctt gca aat aag aat gtt aaa gat gct      583
Gln Ile Asp His Tyr Leu Val Leu Ala Asn Lys Asn Val Lys Asp Ala
          130          135          140
atg gct aaa atc caa gca aaa atc cct gga ttg aag cgc aaa gct gaa      631
Met Ala Lys Ile Gln Ala Lys Ile Pro Gly Leu Lys Arg Lys Ala Glu
          145          150          155
tgaaaacgcc caaaataatt agtaggagtt catcttttaa ggggatattc atttgattat      691
acggggggagg gtcaggaag aacgaacctt gacgttgcag tgcagtttca cagatcgttg      751
ttagatcttt attttttagcc atgcactgtt gtgaggaaaa attacctgtc ttgactgcc      811
tgtgttcac atcttaagta ttgtaagctg ctatgtatgg atttaaaccg taatcatatc      871
tttttccat ctatctgagg cactgggtgga ataaaaaacc tgtatatttt actttgttgc      931
agatagtctt gccgcattct ggcaagttgc agagatggtg gagctagaaa aaaaaaaaaac      991
aaa

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<210> 52  
 <211> 412  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 271..399

<400> 52

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gccgctagcg cctcgagcga tgcacctcct ttccaactgg gcaaaccocg cttccagcag      60
acgtccttct atggccgctt caggcacttc ttggatatca tcgaccctcg cacactcttt      120
gtcactgaga gacgtctcag agaggctgtg cagctgctgg aggactataa gcatgggacc      180
ctgcgccccg gggtcaccaa tgaacagctc tggagtgcac agaaaatcaa gcaggctatt      240
ctacatccgg acaccaatga gaagatcttc atg cca ttt aga atg tca ggt tat      294
                               Met Pro Phe Arg Met Ser Gly Tyr
                               1           5
att cct ttt ggg acg cca att gta agt gtt acc ttc aaa gga ttt cct      342
Ile Pro Phe Gly Thr Pro Ile Val Ser Val Thr Phe Lys Gly Phe Pro
    10           15           20
ttt cta aaa aat tat ttt aaa tgt cta act tta tgt tat tgc tca cgg      390
Phe Leu Lys Asn Tyr Phe Lys Cys Leu Thr Leu Cys Tyr Cys Ser Arg
    25           30           35           40
gta ttt gac tgaattgttg att      412
Val Phe Asp

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<210> 53

<211> 597

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 103..252

<221> sig\_peptide

<222> 103..213

<223> Von Heijne matrix

score 3.9

seq PGPSLRLFSGSQA/SV

<221> polyA\_site

<222> 588..597

<400> 53

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gaaagggtcag aggaaggagc tgtgggaagc tcgcagcagg tatcggagct taagccagtg      60
gatttgggggg ccttgggctc cctagccggc tgcggtgtga ga atg gag tgg gca      114
                               Met Glu Trp Ala
                               -35
gga aag cag cgg gac ttt cag gta agg gca gct ccg ggc tgg gat cat      162
Gly Lys Gln Arg Asp Phe Gln Val Arg Ala Ala Pro Gly Trp Asp His
    -30           -25           -20
ttg gcc tcc ttt cct ggc cct tct ctc cgg ctg ttt tct ggg agt cag      210
Leu Ala Ser Phe Pro Gly Pro Ser Leu Arg Leu Phe Ser Gly Ser Gln
    -15           -10           -5
gcg agt gtc tgt agt ctc tgc tcg ggg ttt ggg gct cag gaa      252
Ala Ser Val Cys Ser Leu Cys Ser Gly Phe Gly Ala Gln Glu
    1           5           10
tgatgtcatg ctccaacagt tggattctat tagcttaagg aggagggaaa cagccaattt      312
tcttgacttt gcaaatctag ctgatctcac tcttgctgaa tctgaggtgt ttagacttca      372
ctctaaaaag catcatttta cttttattta gcacaaaggc acaggatatt tttacaggaa      432
gaatctttta tatggaaaaa tctgagttaa catcactccc gtggtgtttg tagttcttac      492
agggaaactc cagtgccttt tgagccgctt gttcgtccta gtgaacactg tctgttttgt      552

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<210> 55  
<211> 703  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 31..231

<221> polyA\_signal  
<222> 769..774

<221> polyA\_site  
<222> 690..703

<400> 55  
ctctggtggc tctgctacgg cggcgcagaa atg agg cag aag cgg aaa gga gat 54  
Met Arg Gln Lys Arg Lys Gly Asp  
1 5  
ctc agc cct gct aag ctg atg atg ctg act ata gga gat gtt att aaa 102  
Leu Ser Pro Ala Lys Leu Met Met Leu Thr Ile Gly Asp Val Ile Lys  
10 15 20  
caa ctg att gaa gcc cac gag cag ggg aaa gac atc gat cta aat aag 150  
Gln Leu Ile Glu Ala His Glu Gln Gly Lys Asp Ile Asp Leu Asn Lys  
25 30 35 40  
gtg aga acc aag aca gct gcc aaa tat ggc ctt tct gcc cag ccc cgc 198  
Val Arg Thr Lys Thr Ala Ala Lys Tyr Gly Leu Ser Ala Gln Pro Arg  
45 50 55  
ctg gtg gat atc att gct gcc gtc cct cct gag tagctgggat tacaggcacc 251  
Leu Val Asp Ile Ile Ala Ala Val Pro Pro Glu  
60 65  
cgccgctgcc aatttttgta ttttttagtag ggatgggggt ttcaccatat tggtcaggct 311  
ggtctcgaac tctgacctc aggtgatcaa cccaccttgg cctccctaaa tgccgggatt 371  
acaggcatga gccacgctc cgggcctttg attttttaag gtggattttg gttgttataa 431  
atggagaaag gtaagagttc aagttcaacc cgtgtgtgaa agcaaaacaa tggaaaacag 491  
gattggcttc ttcaaaggct cctctttag aactgcctct ttgaaatttc gaggtaatct 551  
actttggaga ctctgcctgg agagggtcag ttcttaagtt aaaagcatcg cttaaccttg 611  
gctcctgtgg cattttacaa aggttttaag gaattgattc ctctgaaagg gcctgaaaat 671  
aaaaagtctt taacatacaa aaaaaaaaaa aa 703

<210> 56  
<211> 725  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 305..565

<221> polyA\_signal  
<222> 694..699

<221> polyA\_site  
<222> 713..725

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<400> 56
ctcacggtgg tgaaggtcac agggttgcag cactcccagt agaccaggag ctccgggagg      60
cagggccggc cccacgtcct ctgcgcacca ccctgagttg gatcctctgt gcgccacccc      120
tgagttggat ccagggctag ctgctgttga cctccccact cccacgctgc cctcctgcct      180
gcagccatga cgcctctgct caccctgacg ctggtgggtcc tcatgggctt acctctggcc      240
caggccttgg actgccacgt gtgaggacta caaatccctc caggatatca ttgccatcct      300
gggt atg gat gaa ctt tct gag gaa gac aag ttg acc gtg tcc cgt gca      349
Met Asp Glu Leu Ser Glu Glu Asp Lys Leu Thr Val Ser Arg Ala
      1          5          10          15
cgg aaa ata cag cgt ttc ttg tct cag cca ttc cag gtt gct gag gtc      397
Arg Lys Ile Gln Arg Phe Leu Ser Gln Pro Phe Gln Val Ala Glu Val
      20          25          30
ttc aca ggt cat atg ggg aag ctg gta ccc ctg aag gag acc atc aaa      445
Phe Thr Gly His Met Gly Lys Leu Val Pro Leu Lys Glu Thr Ile Lys
      35          40          45
gga ttc cag cag att ttg gca ggt gaa tat gac cat ctc cca gaa cag      493
Gly Phe Gln Gln Ile Leu Ala Gly Glu Tyr Asp His Leu Pro Glu Gln
      50          55          60
gcc ttc tat atg gtg gga ccc att gaa gaa gct gtg gca aaa gct gat      541
Ala Phe Tyr Met Val Gly Pro Ile Glu Glu Ala Val Ala Lys Ala Asp
      65          70          75
aag ctg gct gaa gag cat tca tcg tgaggggtct ttgtcctctg tactgtctct      595
Lys Leu Ala Glu Glu His Ser Ser
      80          85
ctccttgccc ctaacccaaa aagcttcatt tttctgtgta ggctgcacaa gagccttgat      655
tgaagatata ttctttctga acagtattta aggtttccaa taaagtgtac acccctcaaa      715
aaaaaaaaaa      725

<210> 57
<211> 1705
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 124..873

<221> sig_peptide
<222> 124..378
<223> Von Heijne matrix
      score 3.6
      seq HLSVVTLAARKVC/IP

<221> polyA_signal
<222> 1673..1678

<221> polyA_site
<222> 1694..1705

<400> 57
cggaggtgag gagcggcggc cccgcccggg gcgctggagg tcgaagcttc caggtagcgg      60
cccgcagagc ctgaccacagg ctctggacat cctgagccca agtccccac actcagtgca      120
gtg atg agt gcg gaa gtg aag gtg aca ggg cag aac cag gag caa ttt      168
Met Ser Ala Glu Val Lys Val Thr Gly Gln Asn Gln Glu Gln Phe

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	-85					-80					-75					
ctg	ctc	cta	gcc	aag	tgc	gcc	aag	ggg	gca	gcg	ctg	gcc	aca	ctc	atc	216
Leu	Leu	Leu	Ala	Lys	Ser	Ala	Lys	Gly	Ala	Ala	Leu	Ala	Thr	Leu	Ile	
-70					-65					-60					-55	
cat	cag	gtg	ctg	gag	gcc	cct	ggt	gtc	tac	gtg	ttt	gga	gaa	ctg	ctg	264
His	Gln	Val	Leu	Glu	Ala	Pro	Gly	Val	Tyr	Val	Phe	Gly	Glu	Leu	Leu	
				-50					-45					-40		
gac	atg	ccc	aat	gtt	aga	gag	ctg	naa	gcc	cgg	aat	ctt	cct	cca	cta	312
Asp	Met	Pro	Asn	Val	Arg	Glu	Leu	Xaa	Ala	Arg	Asn	Leu	Pro	Pro	Leu	
			-35					-30					-25			
aca	gag	gct	cag	aag	aat	aag	ctt	cga	cac	ctc	tca	gtt	gtc	acc	ctg	360
Thr	Glu	Ala	Gln	Lys	Asn	Lys	Leu	Arg	His	Leu	Ser	Val	Val	Thr	Leu	
		-20					-15					-10				
gct	gct	aaa	gta	aag	tgt	atc	cca	tat	gca	gtg	ttg	ctg	gag	gct	ctt	408
Ala	Ala	Lys	Val	Lys	Cys	Ile	Pro	Tyr	Ala	Val	Leu	Leu	Glu	Ala	Leu	
-5						1				5					10	
gcc	ctg	cgt	aat	gtg	cgg	cag	ctg	gaa	gac	ctt	gtg	att	gag	gct	gtg	456
Ala	Leu	Arg	Asn	Val	Arg	Gln	Leu	Glu	Asp	Leu	Val	Ile	Glu	Ala	Val	
			15						20					25		
tat	gct	gac	gtg	ctt	cgt	ggc	tcc	ctg	gac	cag	cgc	aac	cag	cgg	ctc	504
Tyr	Ala	Asp	Val	Leu	Arg	Gly	Ser	Leu	Asp	Gln	Arg	Asn	Gln	Arg	Leu	
			30					35					40			
gag	gtt	gac	tac	agc	atc	ggg	cgg	gac	atc	cag	cgc	cag	gac	ctc	agt	552
Glu	Val	Asp	Tyr	Ser	Ile	Gly	Arg	Asp	Ile	Gln	Arg	Gln	Asp	Leu	Ser	
		45					50					55				
gcc	att	gcc	cga	acc	ctg	cag	gaa	tgg	tgt	gtg	ggc	tgt	gag	gtc	gtg	600
Ala	Ile	Ala	Arg	Thr	Leu	Gln	Glu	Trp	Cys	Val	Gly	Cys	Glu	Val	Val	
	60					65					70					
ctg	tca	ggc	att	gag	gag	cag	gtg	agc	cgt	gcc	aac	caa	cac	aag	gag	648
Leu	Ser	Gly	Ile	Glu	Glu	Gln	Val	Ser	Arg	Ala	Asn	Gln	His	Lys	Glu	
75					80					85					90	
cag	cag	ctg	ggc	ctg	aag	cag	cag	att	gag	agt	gag	gtt	gcc	aac	ctt	696
Gln	Gln	Leu	Gly	Leu	Lys	Gln	Gln	Ile	Glu	Ser	Glu	Val	Ala	Asn	Leu	
			95						100					105		
aaa	aaa	acc	att	aaa	gtt	acg	acg	gca	gca	gca	gcc	gca	gcc	aca	tct	744
Lys	Lys	Thr	Ile	Lys	Val	Thr	Thr	Ala	Ala	Ala	Ala	Ala	Ala	Thr	Ser	
		110						115					120			
cag	gac	cct	gag	caa	cac	ctg	act	gag	ctg	agg	gaa	cca	gct	cct	ggc	792
Gln	Asp	Pro	Glu	Gln	His	Leu	Thr	Glu	Leu	Arg	Glu	Pro	Ala	Pro	Gly	
		125					130					135				
acc	aac	cag	cgc	cag	ccc	agc										

```
tggcaatcct ggaagggttt aatctccttt tgtgagtttg gtggggaagg gaagggtata 1613
tagattatat taaaaaaaaa aggtatatata tgcatatatc tatatataat atgacgcaga 1673
aataaatcta tgagaaatcc aaaaaaaaaa aa 1705
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<210> 58  
<211> 1069  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 135..206

<221> polyA\_signal  
<222> 850..855

<221> polyA\_site  
<222> 1056..1069

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<400> 58
ccccactccgc tctcacgact aagctctcac gattaaggca cgctgcctc gattgtccag 60
ccctctgccag aagaaagctt agcagccagc gcctcagtag agacctaagg gcgctgaatg 120
agtgaggaaag ggaa atg ccg acc aat tgc gct gcg gcg ggc tgt gcc act 170
Met Pro Thr Asn Cys Ala Ala Ala Gly Cys Ala Thr
1 5 10
acc tac aac aag cac att aac atc agc ttc cac agg taacctgggc 216
Thr Tyr Asn Lys His Ile Asn Ile Ser Phe His Arg
15 20
agggagtggg ggtgacggaa actggagttc ctattgtggc tatcgcttgt gtggaaggaa 276
caggaggatt ctgctaattc taataacttt cccagctggg agcagggaag catcgatatgt 336
cctttgtgtt tctcaaactc gcccaattgt tctctgcttt cggggaagct ttactcattt 396
tcttaaaagaa atccaagtac tgtttggtca ttacccttta gtaaaaaaaaa gtaacaggag 456
gatatcgtaa ttttctactg ttttattcct ctgtttagacc gggccttgac atgaatgacg 516
ccgtaaggga gaaagagatc ttoccaatca gcaatcaccg taaaagcctg ctgtgttccc 576
gttaaaatta ggaaattctc actagatgaa ttgacatggg aggcatttag atttctaata 636
gtcacatagt aattctgcgg aggaattgag tcatctttga tagccatgga attaagcgat 696
gttaattaaa gtgcaaacga taacctttct gttcttacta gaatagagta ataaaaagaa 756
cctaggtttt cttttgtttg ctggaagaaa aatcaaaatt ctttagttct gtcaaaccag 816
aactcttgaa agcactttga acaatgcctg gaaaataaca ggtactctgt aaatgtttac 876
cttctctgca agtgccctgcc acgtgcccga agaaaagaca cattaaaaag ttaagtgaca 936
ccagtcctga ttttatatat tttatatacc taacaacgta tatgttagta ttagagaaatt 996
atatccttga cctttttccc tacctattac gaactgtact tttattaaaa gctgccacta 1056
aaaaaaaaaa aaa 1069
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<210> 59  
<211> 1084  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 135..818

<221> polyA\_signal

<222> 909..914

<221> polyA\_site

<222> 1071..1084

<400> 59

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cccactccgc tctcagcact aagctctcac gattaaggca cgctgcctc gattgtccag      60
cctctgccag aagaaagctt agcagccagc gcctcagtag aggcctaagg gcgctgaatg      120
agtgggaaag ggaa atg ccg acc aat tgc gct gcg gcg ggc tgt gcc act      170
                Met Pro Thr Asn Cys Ala Ala Ala Gly Cys Ala Thr
                1          5          10
acc tac aac aag cac att aac atc agc ttc cac agg ttt cct ttg gat      218
Thr Tyr Asn Lys His Ile Asn Ile Ser Phe His Arg Phe Pro Leu Asp
                15          20          25
cct aaa aga aga aaa gaa tgg gtt cgc ctg gtt agg cgc aaa aat ttt      266
Pro Lys Arg Arg Lys Glu Trp Val Arg Leu Val Arg Arg Lys Asn Phe
                30          35          40
gtg cca gga aaa cac act ttt ctt tgt tca aag cac ttt gaa gcc tcc      314
Val Pro Gly Lys His Thr Phe Leu Cys Ser Lys His Phe Glu Ala Ser
                45          50          55          60
tgt ttt gac cta aca gga caa act cga cga ctt aaa atg gat gct gtt      362
Cys Phe Asp Leu Thr Gly Gln Thr Arg Arg Leu Lys Met Asp Ala Val
                65          70          75
cca acc att ttt gat ttt tgt acc cat ata aag tct atg aaa ctc aag      410
Pro Thr Ile Phe Asp Phe Cys Thr His Ile Lys Ser Met Lys Leu Lys
                80          85          90
tca agg aat ctt ttg aag aaa aac aac agt tgt tct cca gct gga cca      458
Ser Arg Asn Leu Leu Lys Lys Asn Asn Ser Cys Ser Pro Ala Gly Pro
                95          100          105
tct agt tta aaa tca aac att agt agt cag caa gta cta ctt gaa cac      506
Ser Ser Leu Lys Ser Asn Ile Ser Ser Gln Gln Val Leu Leu Glu His
                110          115          120
agc tat gcc ttt agg aat cct atg gag gca aaa aag agg atc att aaa      554
Ser Tyr Ala Phe Arg Asn Pro Met Glu Ala Lys Lys Arg Ile Ile Lys
                125          130          135          140
ctg gaa aaa gaa ata gca agc tta aga aga aaa atg aaa act tgc cta      602
Leu Glu Lys Glu Ile Ala Ser Leu Arg Arg Lys Met Lys Thr Cys Leu
                145          150          155
caa aag gaa cgc aga gca act cga aga tgg atc aaa gcc atg tgt ttg      650
Gln Lys Glu Arg Arg Ala Thr Arg Arg Trp Ile Lys Ala Met Cys Leu
                160          165          170
gta aag aat tta gaa gca aat agt gta tta cct aaa ggt aca tca gaa      698
Val Lys Asn Leu Glu Ala Asn Ser Val Leu Pro Lys Gly Thr Ser Glu
                175          180          185
cac atg tta cca act gcc tta agc agt ctt cct ttg gaa gat ttt aag      746
His Met Leu Pro Thr Ala Leu Ser Ser Leu Pro Leu Glu Asp Phe Lys
                190          195          200
atc ctt gaa caa gat caa caa gat aaa aca ctg cta agt cta aat cta      794
Ile Leu Glu Gln Asp Gln Gln Asp Lys Thr Leu Leu Ser Leu Asn Leu
                205          210          215          220
aaa cag acc aag agt acc ttc att taaatttagc ttgcacagag cttgatgcct      848
Lys Gln Thr Lys Ser Thr Phe Ile
                225
atccttcatt cttttcagaa gtaaagataa ttatggcact tatgccaaaa ttcattattt      908
aataaagttt tacttgaagt aacattactg aatttgtgaa gacttgatta caaaagaata      968
aaaaacttca tatggaaatt ttatttgaaa atgagtggaa gcgcottaca ttagaattac      1028

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ggacttaaaa attttgctaa taaattgtgt gtttgaaagg tgaaaaaaa aaaaaa 1084

<210> 60  
<211> 419  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 33..290

<221> sig\_peptide  
<222> 33..92  
<223> Von Heijne matrix  
score 5.4  
seq WFFVHSSALGLVLA/PP

<400> 60  
aatggtaggc cttcatgtga gccagttact ac atg aat ctt cat ttc cca cag 53  
Met Asn Leu His Phe Pro Gln  
-20 -15  
tgg ttt gtt cat tca tca gcg tta ggc ttg gtc ctg gct cca cct ttc 101  
Trp Phe Val His Ser Ser Ala Leu Gly Leu Val Leu Ala Pro Pro Phe  
-10 -5 1  
tcc tct ccg ggc act gac ccc acc ttt ccg tgt att tac tgt agg cta 149  
Ser Ser Pro Gly Thr Asp Pro Thr Phe Pro Cys Ile Tyr Cys Arg Leu  
5 10 15  
tta aat atg atc atg acc cgc ctt gca ttt tca ttc atc acc tgt tta 197  
Leu Asn Met Ile Met Thr Arg Leu Ala Phe Ser Phe Ile Thr Cys Leu  
20 25 30 35  
tgc cca aat tta aag gaa gtt tgt ctc att ttg cca gaa aaa aat tgt 245  
Cys Pro Asn Leu Lys Glu Val Cys Leu Ile Leu Pro Glu Lys Asn Cys  
40 45 50  
aat agt cgg cac gct gga ttt gta ggg cca gca aaa ttg cgg cag 290  
Asn Ser Arg His Ala Gly Phe Val Gly Pro Ala Lys Leu Arg Gln  
55 60 65  
tgaaactagt ttcacttcta aagcccttca tttccacaa ggttaagctc tcgaaacccc 350  
atttgatcct tggttcctat ttogatcctc ctttgggaatc tgaaaatcgg tctccatgtt 410  
gtatgcaaa 419

<210> 61  
<211> 682  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 485..616

<221> polyA\_site  
<222> 669..682

<400> 61

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ctcctttctc attccttata ttgcgtgttt ttaccttttt ttcataacta agtttttgag    60
gaagtttagtg ttcttttcaa agaaccggtt cgaaatgtac ttttctttgc tactttttgt    120
tattttattg atcacatctt taatcttttg ttctctatac gtggcctgtt ttgatttatt    180
ttactattct tgctttctaa ggtaagtatt ttgttggtga gtgctttatt tttttcatct    240
ttcttcttga ataataatga catttttagg ttataaatTT tctcttggtg ctcagtttgc    300
ctcatTAatt ttggcagtaa gcattctcct tttattgctt tctatgtagt ctttaatttt    360
gcttttaact tcttctttga tctaaggatt acctacttgt taatttccaa atattatctt    420
atctatctat ctatctatct atctatctat ctatctatct acctatgtga gacgaagtct    480
ggct atg tcg ccg agg ctg gag tgc agt ggt gca atc ttg gct cac tgc    529
Met Ser Pro Arg Leu Glu Cys Ser Gly Ala Ile Leu Ala His Cys
1         5         10        15
aac ccc cgc ctc cca ggt tca agt tat tct cct gcc tca gct act tgg    577
Asn Pro Arg Leu Pro Gly Ser Ser Tyr Ser Pro Ala Ser Ala Thr Trp
20        25        30
gtg aga gga tcc ctt gag ccg ggg agg ttg agg ctg cag tgagccataa    626
Val Arg Gly Ser Leu Glu Pro Gly Arg Leu Arg Leu Gln
35        40
ccactactct ccagcctgga taacaaaagt gagactctga ccaaaaaaaaa aaaaaa    682

```

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<210> 62
<211> 1191
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 54..995

<221> sig_peptide
<222> 54..227
<223> Von Heijne matrix
      score 4.1
      seq LVHHCPTWQWATG/EE

<221> polyA_signal
<222> 1130..1135

<221> polyA_site
<222> 1181..1191

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<400> 62
cacggctgca ctttccatcc cgtcgcgggg ccggccgcta ctccggcccc agg atg    56
Met
cag aat gtg att aat act gtg aag gga aag gca ctg gaa gtg gct gag    104
Gln Asn Val Ile Asn Thr Val Lys Gly Lys Ala Leu Glu Val Ala Glu
-55 -50 -45
tac ctg acc ccg gtc ctc aag gaa tca aag ttt agg gaa aca ggt gta    152
Tyr Leu Thr Pro Val Leu Lys Glu Ser Lys Phe Arg Glu Thr Gly Val
-40 -35 -30
att acc cca gaa gag ttt gtg gca gct gga gat cac cta gtc cac cac    200
Ile Thr Pro Glu Glu Phe Val Ala Ala Gly Asp His Leu Val His His
-25 -20 -15 -10
tgt cca aca tgg caa tgg gct aca ggg gaa gaa ttg aaa gtg aag gca    248
Cys Pro Thr Trp Gln Trp Ala Thr Gly Glu Glu Leu Lys Val Lys Ala
-5 1 5

```

tac cta cca aca ggc aaa caa ttt ttg gta acc aaa aat gtg ccg tgc	296
Tyr Leu Pro Thr Gly Lys Gln Phe Leu Val Thr Lys Asn Val Pro Cys	
10 15 20	
tat aag cgg tgc aaa cag atg gaa tat tca gat gaa ttg gaa gct atc	344
Tyr Lys Arg Cys Lys Gln Met Glu Tyr Ser Asp Glu Leu Glu Ala Ile	
25 30 35	
att gaa gaa gat gat ggt gat ggc gga tgg gta gat aca tat cac aac	392
Ile Glu Glu Asp Asp Gly Asp Gly Gly Trp Val Asp Thr Tyr His Asn	
40 45 50 55	
aca ggt att aca gga ata acg gaa gcc gtt aaa gag atc aca ctg gaa	440
Thr Gly Ile Thr Gly Ile Thr Glu Ala Val Lys Glu Ile Thr Leu Glu	
60 65 70	
aat aag gac aat ata agg ctt caa gat tgc tca gca cta tgt gaa gag	488
Asn Lys Asp Asn Ile Arg Leu Gln Asp Cys Ser Ala Leu Cys Glu Glu	
75 80 85	
gaa gaa gat gaa gat gaa gga gaa gct gca gat atg gaa gaa tat gaa	536
Glu Glu Asp Glu Asp Glu Gly Glu Ala Ala Asp Met Glu Glu Tyr Glu	
90 95 100	
gag agt gga ttg ttg gaa aca gat gag gct acc cta gat aca agg aaa	584
Glu Ser Gly Leu Leu Glu Thr Asp Glu Ala Thr Leu Asp Thr Arg Lys	
105 110 115	
ata gta gaa gct tgt aaa gcc aaa act gat gct ggc ggt gaa gat gct	632
Ile Val Glu Ala Cys Lys Ala Lys Thr Asp Ala Gly Gly Glu Asp Ala	
120 125 130 135	
att ttg caa acc aga act tat gac ctt tac atc act tat gat aaa tat	680
Ile Leu Gln Thr Arg Thr Tyr Asp Leu Tyr Ile Thr Tyr Asp Lys Tyr	
140 145 150	
tac cag act cca cga tta tgg ttg ttt ggc tat gat gag caa cgg cag	728
Tyr Gln Thr Pro Arg Leu Trp Leu Phe Gly Tyr Asp Glu Gln Arg Gln	
155 160 165	
cct tta aca gtt gag cac atg tat gaa gac atc agt cag gat cat gtg	776
Pro Leu Thr Val Glu His Met Tyr Glu Asp Ile Ser Gln Asp His Val	
170 175 180	
aag aaa aca gtg acc att gaa aat cat cct cat ctg cca cca cct ccc	824
Lys Lys Thr Val Thr Ile Glu Asn His Pro His Leu Pro Pro Pro Pro	
185 190 195	
atg tgt tca gtt cac cca tgc agg cat gct gag gtg atg aag aaa atc	872
Met Cys Ser Val His Pro Cys Arg His Ala Glu Val Met Lys Lys Ile	
200 205 210 215	
att gag act gtt gca gaa gga ggg gga gaa ctt gga gtt cat atg tat	920
Ile Glu Thr Val Ala Glu Gly Gly Gly Glu Leu Gly Val His Met Tyr	
220 225 230	
ctt ctt att ttc ttg aaa ttt gta caa gct gtc att cca aca ata gaa	968
Leu Leu Ile Phe Leu Lys Phe Val Gln Ala Val Ile Pro Thr Ile Glu	
235 240 245	
tat gac tac aca aga cac ttc aca atg taatgaagag agcataaaat	1015
Tyr Asp Tyr Thr Arg His Phe Thr Met	
250 255	
ctatcctaatt tattggttct gattttttaaa gaattaaccc atagatgtga ccattgacca	1075
tattcatcaa tatatacagt ttctctaata agggacttat atgttttatgc attaaataaa	1135
aatatgttcc actaccagcc ttacttgttt aataaaaaatc agtgcaaaaa aaaaaa	1191

<210> 63  
 <211> 1008  
 <212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 657..923

<221> sig\_peptide

<222> 657..896

<223> Von Heijne matrix

score 3.5

seq RGLLSACAPWGDG/ST

<221> polyA\_signal

<222> 957..962

<221> polyA\_site

<222> 974..1008

<400> 63

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ntcgnatgtg gcacaaaacc cctctgctgg ctcatgtgtg caactgagac tgtcagagca      60
tggctagctc tgggggccag ctctgctggg tgggggctag agaggaagca gggagtatct    120
gcacacagga tgcctgcgct cagggtggtg cagaagtcag tgcccaggcc cccccacaca    180
gtccccaag gtccggcctc cccagcgcgg ggctcctcgt ttgaggggag gtgacttccc    240
tcccagcagg ctcttggaaca cagtaagctt cccagccctt gcctgagcag cttttcctcc    300
ttgccctgtt cccacacctc cggctccagt ccaggggagct cccagggaag tggtcgaccc    360
ctccagtggc tgggccaactc tgctagagtc catccgccaa gctgggggca tcggcaaggc    420
caagctgcgc agcatgaagg agcgaagct ggagaagaag aagcagaagg agcaggagca    480
agtgagagcc acgagccaag gtgggcactt gatgtcggat ctcttcaaca agctgggtcat    540
gaggcgcaag ggcattctctg ggaaagaacc tggggctggt gaggggcccg gaggagcctt    600
tgcccgctg tcagactcca tccctcctct gccgccaccg cagcagccac aggtag atg    659
                                         Met
                                         -80
agg aca agg acg act ggg aat cct agg ggg ctc cat gac acc ttc ccc      707
Arg Thr Arg Thr Thr Gly Asn Pro Arg Gly Leu His Asp Thr Phe Pro
          -75                      -70                      -65
cgc aga ccc aga ctt ggc cgt tgc tct gac atg gac aca gcc agg aca      755
Arg Arg Pro Arg Leu Gly Arg Cys Ser Asp Met Asp Thr Ala Arg Thr
          -60                      -55                      -50
agc tgc tca gac ctg ctt ccc tgg gag ggg gtg acg gaa cca gca ctg      803
Ser Cys Ser Asp Leu Leu Pro Trp Glu Gly Val Thr Glu Pro Ala Leu
          -45                      -40                      -35
tgt gga gac cag ctt caa gga acg gaa ggc tgg ctt gag gcc aca cag      851
Cys Gly Asp Gln Leu Gln Gly Thr Glu Gly Trp Leu Glu Ala Thr Gln
          -30                      -25                      -20
ctg ggg cgg gga ctt ctg tct gcc tgt gct cca tgg ggg gac ggc tcc      899
Leu Gly Arg Gly Leu Leu Ser Ala Cys Ala Pro Trp Gly Asp Gly Ser
          -15                      -10                      -5                      1
acc cag cct gtg cca ctg tgt tct taagaggctt ccagagaaaa cggcacacca      953
Thr Gln Pro Val Pro Leu Cys Ser
          5
atcaataaag aactgagcag aaaaaaaaaa aaaaaaaaaa aaaaaaaaaa aaaaan    1008
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<210> 64

<211> 568

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 18..311

<221> sig\_peptide

<222> 18..62

<223> Von Heijne matrix

score 8.4

seq AMWLLCVALAVLA/WG

<400> 64

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agtgctgctt acccatc atg gaa gca atg tgg ctc ctg tgt gtg gcg ttg      50
                Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu
                -15                -10                -5
gcg gtc ttg gca tgg ggc ttc ctc tgg gtt tgg gac tcc tca gaa cga      98
Ala Val Leu Ala Trp Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg
                1                5                10
atg aag agt cgg gag cag gga gga cgg ctg gga gcc gaa agc cgg acc      146
Met Lys Ser Arg Glu Gln Gly Gly Arg Leu Gly Ala Glu Ser Arg Thr
                15                20                25
ctg ctg gtc ata gcg cac cct gac gat gaa gcc atg ttt ttt gct ccc      194
Leu Leu Val Ile Ala His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro
                30                35                40
aca gtg cta ggc ttg gcc cgc cta agg cac tgg gtg tac ctg ctt tgc      242
Thr Val Leu Gly Leu Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys
                45                50                55                60
ttc tct gca gtt ttc cgt agg gag cta agt gaa tac acc gaa ggt ctt      290
Phe Ser Ala Val Phe Arg Arg Glu Leu Ser Glu Tyr Thr Glu Gly Leu
                65                70                75
acc tct gaa ccc ctc aca gcc tagggacagg agcggccggc ttacctggtg      341
Thr Ser Glu Pro Leu Thr Ala
                80
ggttggggga cgctggcagc tcgcgtacta cgccagcagg attgaggagc agagaaacag      401
ttgcagttgg ttgtattcag tacctgcatt tccgttggga actccacctg tacttggtat      461
tctgtggaac tttttttatt tgtagaagga gcaagaatat tgaccttact atatagcaca      521
cgaaacaatc tatgctgtat cgtgcctgct caatccttaa agttaac      568

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<210> 65

<211> 538

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 151..426

<221> sig\_peptide

<222> 151..258

<223> Von Heijne matrix

score 5.2

seq KVALAGLLGFGLG/KV

<221> polyA\_signal

<222> 505..510

<221> polyA\_site

<222> 527..538

<400> 65

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cactgggtca aggagtaagc agaggataaa caactggaag gagagcaagc acaaagtcac      60
catggcttca gcgtctgctc gtggaaacca agataaagat gcccatcttc caccaccaag      120
caagcagctc tgcctttttc tcttgtaagc atg ctt gtc acc cag gga cta gtc      174
                               Met Leu Val Thr Gln Gly Leu Val
                               -35                               -30
tac caa ggt tat ttg gca gct aat tct aga ttt gga tca ttg ccc aaa      222
Tyr Gln Gly Tyr Leu Ala Ala Asn Ser Arg Phe Gly Ser Leu Pro Lys
                               -25                               -20                               -15
gtt gca ctt gct ggt ctc ttg gga ttt ggc ctt gga aag gta tca tac      270
Val Ala Leu Ala Gly Leu Leu Gly Phe Gly Leu Gly Lys Val Ser Tyr
                               -10                               -5                               1
ata gga gta tgc cag agt aaa ttc cat ttt ttt gaa gat cag ctc cgt      318
Ile Gly Val Cys Gln Ser Lys Phe His Phe Phe Glu Asp Gln Leu Arg
5                               10                               15                               20
ggg gct ggt ttt ggt cca cag cat aac agg cac tgc ctc ctt acc tgt      366
Gly Ala Gly Phe Gly Pro Gln His Asn Arg His Cys Leu Leu Thr Cys
                               25                               30                               35
gag gaa tgc aaa ata aag cat gga tta agt gag aag gga gac tct cag      414
Glu Glu Cys Lys Ile Lys His Gly Leu Ser Glu Lys Gly Asp Ser Gln
                               40                               45                               50
cct tca gct tcc taaattctgt gtctgtgact ttcgaagttt tttaaacttc      466
Pro Ser Ala Ser
                               55
tgaatttgta cacatttaaa atttcaagtg tacttttaaaa taaaatactt ctaatggaac      526
aaaaaaaaaa aa      538

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<210> 66

<211> 1747

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 10..1062

<221> sig\_peptide

<222> 10..57

<223> Von Heijne matrix

score 4.9

seq FIYLAHFTLCSG/WS

<221> polyA\_signal

<222> 1710..1715

<221> polyA\_site

<222> 1735..1747

<400> 66

```

gcctcacca atg gtt ccc ttc atc tat ctg caa gcc cac ttt aca ctc tgt      51

```

Met Val Pro Phe Ile Tyr Leu Gln Ala His Phe Thr Leu Cys																
-15					-10					-5						
tct	ggg	tgg	tcc	agc	aca	tac	cgg	gac	ctc	cgg	aag	ggt	gtg	tat	gtg	99
Ser	Gly	Trp	Ser	Ser	Thr	Tyr	Arg	Asp	Leu	Arg	Lys	Gly	Val	Tyr	Val	
	1					5				10						
ccc	tac	acc	cag	ggc	aag	tgg	gaa	ggg	gag	ctg	ggc	acc	gac	ctg	gta	147
Pro	Tyr	Thr	Gln	Gly	Lys	Trp	Glu	Gly	Glu	Leu	Gly	Thr	Asp	Leu	Val	
15					20					25					30	
agc	atc	ccc	cat	ggc	ccc	aac	gtc	act	gtg	cgt	gcc	aac	att	gct	gcc	195
Ser	Ile	Pro	His	Gly	Pro	Asn	Val	Thr	Val	Arg	Ala	Asn	Ile	Ala	Ala	
				35					40					45		
atc	act	gaa	tca	gac	aag	ttc	ttc	atc	aac	ggc	tcc	aac	tgg	gaa	ggc	243
Ile	Thr	Glu	Ser	Asp	Lys	Phe	Phe	Ile	Asn	Gly	Ser	Asn	Trp	Glu	Gly	
			50					55					60			
atc	ctg	ggg	ctg	gcc	tat	gct	gag	att	gcc	agg	cct	gac	gac	tcc	ccg	291
Ile	Leu	Gly	Leu	Ala	Tyr	Ala	Glu	Ile	Ala	Arg	Pro	Asp	Asp	Ser	Pro	
			65				70					75				
gag	cct	ttc	ttt	gac	tct	ctg	gta	aag	cag	acc	cac	gtt	ccc	aac	ctc	339
Glu	Pro	Phe	Phe	Asp	Ser	Leu	Val	Lys	Gln	Thr	His	Val	Pro	Asn	Leu	
	80					85					90					
ttc	tcc	ctg	cag	ctt	tgt	ggt	gct	ggc	ttc	ccc	ctc	aac	cag	tct	gaa	387
Phe	Ser	Leu	Gln	Leu	Cys	Gly	Ala	Gly	Phe	Pro	Leu	Asn	Gln	Ser	Glu	
95					100				105						110	
gtg	ctg	gcc	tct	gtc	gga	ggg	agc	atg	atc	att	gga	ggt	atc	gac	cac	435
Val	Leu	Ala	Ser	Val	Gly	Gly	Ser	Met	Ile	Ile	Gly	Gly	Ile	Asp	His	
				115					120					125		
tcg	ctg	tac	aca	ggc	agt	ctc	tgg	tat	aca	ccc	atc	cgg	cgg	gag	tgg	483
Ser	Leu	Tyr	Thr	Gly	Ser	Leu	Trp	Tyr	Thr	Pro	Ile	Arg	Arg	Glu	Trp	
			130					135					140			
tat	tat	gag	gtg	atc	att	gtg	cgg	gtg	gag	atc	aat	gga	cag	gat	ctg	531
Tyr	Tyr	Glu	Val	Ile	Ile	Val	Arg	Val	Glu	Ile	Asn	Gly	Gln	Asp	Leu	
			145			150					155					
aaa	atg	gac	tgc	aag	gag	tac	aac	tat	gac	aag	agc	att	gtg	gac	agt	579
Lys	Met	Asp	Cys	Lys	Glu	Tyr	Asn	Tyr	Asp	Lys	Ser	Ile	Val	Asp	Ser	
	160					165					170					
ggc	acc	acc	aac	ctt	cgt	ttg	ccc	aag	aaa	gtg	ttt	gaa	gct	gca	gtc	627
Gly	Thr	Thr	Asn	Leu	Arg	Leu	Pro	Lys	Lys	Val	Phe	Glu	Ala	Ala	Val	
175					180					185					190	
aaa	tcc	atc	aag	gca	gcc	tcc	tcc	acg	gag	aag	ttc	cct	gac	ggt	ttc	675
Lys	Ser	Ile	Lys	Ala	Ala	Ser	Ser	Thr	Glu	Lys	Phe	Pro	Asp	Gly	Phe	
			195						200					205		
tgg	cta	gga	gag	cag	ctg	gtg	tgc	tgg	caa	gca	ggc	acc	acc	cct	tgg	723
Trp	Leu	Gly	Glu	Gln	Leu	Val	Cys	Trp	Gln	Ala	Gly	Thr	Thr	Pro	Trp	
			210					215					220			
aac	att	ttc	cca	gtc	atc	tca	ctc	tac	cta	atg	ggt	gag	gtt	acc	aac	771
Asn	Ile	Phe	Pro	Val	Ile	Ser	Leu	Tyr	Leu	Met	Gly	Glu	Val	Thr	Asn	
			225			230						235				
cag	tcc	ttc	cgc	atc	acc	atc	ctt	ccg	cag	caa	tac	ctg	cgg	cca	gtg	819
Gln	Ser	Phe	Arg	Ile	Thr	Ile	Leu	Pro	Gln	Gln	Tyr	Leu	Arg	Pro	Val	
			240			245					250					
gaa	gat	gtg	gcc	acg	tcc	caa	gac	gac	tgt	tac	aag	ttt	gcc	atc	tca	867
Glu	Asp	Val	Ala	Thr	Ser	Gln	Asp	Asp	Cys	Tyr	Lys	Phe	Ala	Ile	Ser	
255					260					265					270	
cag	tca	tcc	acg	ggc	act	gtt	atg	gga	gct	gtt	atc	atg	gag	ggc	ttc	915
Gln	Ser	Ser	Thr	Gly	Thr	Val	Met	Gly	Ala	Val	Ile	Met	Glu	Gly	Phe	
			275						280					285		
tac	gtt	gtc	ttt	gat	cgg	gcc	cga	aaa	cga	att	ggc	ttt	gct	gtc	agc	963

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Tyr Val Val Phe Asp Arg Ala Arg Lys Arg Ile Gly Phe Ala Val Ser
      290                      295                      300
gct tgc cat gtg cac gat gag ttc agg acg gca gcg gtg gaa ggc ccn      1011
Ala Cys His Val His Asp Glu Phe Arg Thr Ala Ala Val Glu Gly Pro
      305                      310                      315
ttt tgt cac ctt gga cat gga aga ctg tgg cta caa cat tcc aca gac      1059
Phe Cys His Leu Gly His Gly Arg Leu Trp Leu Gln His Ser Thr Asp
      320                      325                      330
aga tgagtcaacc ctcatgacca tagcctatgt catggctgcc atctgcgccc      1112
Arg
335
tcttcatgct gccactctgc ctcatggtgt gtcagtggcg ctgcctccgc tgcctgcgcc      1172
agcagcatga tgactttgct gatgacatct ccctgctgaa gtgaggaggc ccatgggcag      1232
aagataggga ttcccttgga ccacacctcc gtggttcaact ttggtcacaa gtaggagaca      1292
cagatggcac ctgtggccag agcacctcag gaccctcccc acccaccaaa tgcctctgcc      1352
ttgatggaga aggaaaaggc tggcaagggtg gggtccagggt actgtacctg taggagacag      1412
aaaagagaag aaagaagcac tctgctggcg ggaatactct tggtcacctc aaatttaagt      1472
cgggaaattc tgctgcttga aacttcagcc ctgaaccttt gtcaccattc ctttaaattc      1532
tccaacccaa agtattcttc ttttcttagt ttcagaagta ctggcatcac acgcagggtta      1592
ccttggcgtg tgtccctgtg gtaccctggc agagaagaga ccaagcttgt ttccctgctg      1652
gccaaagtca gtaggagagg atgcacagtt tgctatttgc tttagagaca gggactgtat      1712
aaacaagcct aacattgggtg caaaaaaaaa aaaaaa      1747

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<210> 67

<211> 1686

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 78..491

<221> sig\_peptide

<222> 78..218

<223> Von Heijne matrix

score 5.8

seq LMCFGALIGLCAC/IC

<221> polyA\_signal

<222> 1652..1657

<221> polyA\_site

<222> 1673..1686

<400> 67

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ggtatagccc accagaaagg acagagtcac ttgatgtggt cacaaaatgt gtgagtttca      60
cactaactga gcagttc atg gag aaa ttt gtt gat ccc gga aac cac aat      110
      Met Glu Lys Phe Val Asp Pro Gly Asn His Asn
      -45                      -40
agc ggg att gat ctc ctt agg acc tat ctt tgg cgt tgc cag ttc ctt      158
Ser Gly Ile Asp Leu Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu
      -35                      -30                      -25
tta cct ttt gtg agt tta ggt ttg atg tgc ttt ggg gct ttg atc gga      206
Leu Pro Phe Val Ser Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly
      -20                      -15                      -10                      -5

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ctt tgt gct tgc att tgc cga agc tta tat ccc acc att gcc acg ggc      254
Leu Cys Ala Cys Ile Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly
      1              5              10
att ctc cat ctc ctt gca ggt ctg tgt aca ctg ggc tca gta agt tgt      302
Ile Leu His Leu Leu Ala Gly Leu Cys Thr Leu Gly Ser Val Ser Cys
      15              20              25
tat gtt gct gga att gaa cta ctc cac cag aaa cta gag ctc cct gac      350
Tyr Val Ala Gly Ile Glu Leu Leu His Gln Lys Leu Glu Leu Pro Asp
      30              35              40
aat gta tcc ggt gaa ttt gga tgg tcc ttc tgc ctt gct tgt gtc tct      398
Asn Val Ser Gly Glu Phe Gly Trp Ser Phe Cys Leu Ala Cys Val Ser
      45              50              55              60
gct ccc tta cag ttc atg gct tct gct ctc ttc atc tgg gct gct cac      446
Ala Pro Leu Gln Phe Met Ala Ser Ala Leu Phe Ile Trp Ala Ala His
      65              70              75
acc aac cgg aga gag tac acc tta atg aag gca tat cgt gtg gca      491
Thr Asn Arg Arg Glu Tyr Thr Leu Met Lys Ala Tyr Arg Val Ala
      80              85              90
tgagcaagaa actgcctgct ttacaattgc cattttttatt tttttaaat aatactgata      551
ttttcccccac ctctcaattg tttttaattt ttatttgtgg atataccatt ttattatgaa      611
aatctattttt atttatacac attcaccact aaatacacac ttaataccac taaaatttat      671
gtgggtttact ttaagcgatg ccatctttca aataaaactaa tctaggtcta gacagaaaga      731
aatggataga gacttgacac aaatttatga aagaaaattg ggagtaggaa tgtgaccgaa      791
aacaagttgt gctaattgtct gttagacttt tcagtaaaac caaagtaact gtatctgttc      851
aactaaaaac tctatattag tttctttggg aaacctctca tcgtcaaaac tttatgttca      911
ctttgctggtt gtagatagcc agtcaaccag cagtattagt gctgttttca aagatttaag      971
ctctataaaa ttgggaaatt atctaagatc attttcccta agcattgaca catagcttca      1031
tctgaggtga gatatggcag ctgtttgtat ctgcactgtg tctgtctaca aagagtgaaa      1091
aatacagtggt ttacttgaaa ttttaacttt gtaactgcaa gaattccagt tcggccgggc      1151
gaggattagt attattttta actctccgta agattttcag taccaccaa ttgttttgga      1211
ttttttttct ttcctcttca cataccaggg ttattaaaag tgtgctttct ttttacatta      1271
tattacagtt acaaggtaaa attcctcaac tgctatttat ttattccagc ccagtactat      1331
aaagaacggt tcaccataat gacctccag agctgggaaa cctaccacaa gatctaaagt      1391
tctggctgtc cattaacctc caactatggt ctttatttct tgtggttaata tgatgtgcct      1451
ttccttgccct aaatcccttc ctgggtgtga tcaacattat ttaatgtctt ctaattcagt      1511
cattttttat aagtatgtct ataacattg aactttaaaa aacttattta tttattccac      1571
tactgtagca attgacagat taaaaaaatg taacttcata atttcttacc ataacctcaa      1631
tgtctttttt aaaaaataaa attaaaaatg aaaagagacc caaaaaaaaaa aaaaa      1686

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<210> 68  
 <211> 542  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 69..371

<221> sig\_peptide  
 <222> 69..287  
 <223> Von Heijne matrix  
       score 4  
       seq AVGFLEFWIVLTS/WI

<221> polyA\_signal

<222> 510..515

<221> polyA\_site

<222> 530..542

<400> 68

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tggtacttag ggtcaaggct tgggtcttgc cccgcaaacc cttgggaaga cccggcccca    60
gcgcagct atg aac ctg gag cga gtg tcc aat gag gag aaa ttg aac ctg    110
      Met Asn Leu Glu Arg Val Ser Asn Glu Glu Lys Leu Asn Leu
            -70                -65                -60
tgc cgg aag tac tac ctg ggg ggg ttt gct ttc ttg cct ttt ctc tgg    158
Cys Arg Lys Tyr Tyr Leu Gly Gly Phe Ala Phe Leu Pro Phe Leu Trp
            -55                -50                -45
ttg gtc aac atc ttc tgg ttc tac cga gag gcc ttc ctt gtc cca gcc    206
Leu Val Asn Ile Phe Trp Phe Tyr Arg Glu Ala Phe Leu Val Pro Ala
            -40                -35                -30
tac aca gaa cag agc caa atc aaa ggc tat gtc tgg cgc tca gct gtg    254
Tyr Thr Glu Gln Ser Gln Ile Lys Gly Tyr Val Trp Arg Ser Ala Val
            -25                -20                -15
ggc ttc ctc ttc tgg gtg ata gtg ctc acc tcc tgg atc acc atc ttc    302
Gly Phe Leu Phe Trp Val Ile Val Leu Thr Ser Trp Ile Thr Ile Phe
            -10                -5                1                5
cag atc tac cgg ccc cgc tgg ggt gcc ctt ggg gac tac ctc tcc ttc    350
Gln Ile Tyr Arg Pro Arg Trp Gly Ala Leu Gly Asp Tyr Leu Ser Phe
            10                15                20
acc ata ccc ctg ggc acc ccc tgacaacttc tgcacatact ggggccctgc    401
Thr Ile Pro Leu Gly Thr Pro
            25
ttattctccc aggacaggt ccttaaagca gaggagcctg tcctgggagc cccttctcaa    461
actcctaaga cttgtttctca tgtcccaagt tctctgtctga catcccccaa taaaggaccc    521
taacttttcaa aaaaaaaaaa a    542

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<210> 69

<211> 1174

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 2..757

<221> sig\_peptide

<222> 2..205

<223> Von Heijne matrix

score 7.3

seq LRLILSPLPGAQP/QQ

<221> polyA\_site

<222> 1160..1174

<400> 69

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g atg cct gag ggc ccc gag ctg cac ctg gcc agc cag ttt gtg aat gag    49
      Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu
            -65                -60                -55
gcc tgc agg gcg ctg gtg ttc ggc ggc tgc gtg gag aag tcc tct gtc    97

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Ala	Cys	Arg	Ala	Leu	Val	Phe	Gly	Gly	Cys	Val	Glu	Lys	Ser	Ser	Val		
		-50					-45				-40						
agc	cgc	aac	cct	gag	gtg	ccc	ttt	gag	agc	agt	gcc	tac	cgc	atc	tca		145
Ser	Arg	Asn	Pro	Glu	Val	Pro	Phe	Glu	Ser	Ser	Ala	Tyr	Arg	Ile	Ser		
		-35				-30					-25						
gct	tca	gcc	cgc	ggc	aag	gag	ctg	cgc	ctg	ata	ctg	agc	cct	ctg	cct		193
Ala	Ser	Ala	Arg	Gly	Lys	Glu	Leu	Arg	Leu	Ile	Leu	Ser	Pro	Leu	Pro		
		-20			-15					-10				-5			
ggg	gcc	cag	cct	caa	cag	gag	cca	ctg	gcc	ctg	gtc	ttc	cgc	ttc	ggc		241
Gly	Ala	Gln	Pro	Gln	Gln	Glu	Pro	Leu	Ala	Leu	Val	Phe	Arg	Phe	Gly		
			1				5					10					
atg	tcc	ggc	tct	ttt	cag	ctg	gtg	ccc	cgc	gag	gag	ctg	cca	cgc	cat		289
Met	Ser	Gly	Ser	Phe	Gln	Leu	Val	Pro	Arg	Glu	Glu	Leu	Pro	Arg	His		
		15				20					25						
gcc	cac	ctg	cgc	ttt	tac	acg	gcc	ccg	cct	ggc	ccc	cgg	ctc	gcc	cta		337
Ala	His	Leu	Arg	Phe	Tyr	Thr	Ala	Pro	Pro	Gly	Pro	Arg	Leu	Ala	Leu		
		30				35					40						
tgt	ttc	gtg	gac	atc	cgc	cgg	ttc	ggc	cgc	tgg	gac	ctt	ggg	gga	aag		385
Cys	Phe	Val	Asp	Ile	Arg	Arg	Phe	Gly	Arg	Trp	Asp	Leu	Gly	Gly	Lys		
					50					55					60		
tgg	cag	ccg	ggc	cgc	ggg	ccc	tgt	gtc	ttg	cag	gag	tac	cag	cag	ttc		433
Trp	Gln	Pro	Gly	Arg	Gly	Pro	Cys	Val	Leu	Gln	Glu	Tyr	Gln	Gln	Phe		
				65				70				75					
agg	gag	aat	gtg	cta	cga	aac	cta	gcg	gat	aag	gcc	ttt	gac	cgg	ccc		481
Arg	Glu	Asn	Val	Leu	Arg	Asn	Leu	Ala	Asp	Lys	Ala	Phe	Asp	Arg	Pro		
		80						85				90					
atc	tgc	gag	gcc	ctc	ctg	gac	cag	agg	ttc	ttc	aat	ggc	att	ggc	aac		529
Ile	Cys	Glu	Ala	Leu	Leu	Asp	Gln	Arg	Phe	Phe	Asn	Gly	Ile	Gly	Asn		
		95				100					105						
tat	ctg	cgg	gca	gag	atc	ctg	tac	cgg	ctg	aag	atc	ccc	ccc	ttt	gag		577
Tyr	Leu	Arg	Ala	Glu	Ile	Leu	Tyr	Arg	Leu	Lys	Ile	Pro	Pro	Phe	Glu		
		110				115					120						
aag	gcc	cgc	tcg	gtc	ctg	gag	gcc	ctg	cag	cag	cac	agg	ccg	agc	ccg		625
Lys	Ala	Arg	Ser	Val	Leu	Glu	Ala	Leu	Gln	Gln	His	Arg	Pro	Ser	Pro		
		125			130				135						140		
gag	ctg	acc	ctg	agc	cag	aag	ata	agg	acc	aag	ctg	cag	aat	tca	gac		673
Glu	Leu	Thr	Leu	Ser	Gln	Lys	Ile	Arg	Thr	Lys	Leu	Gln	Asn	Ser	Asp		
				145				150						155			
ctg	ctg	gag	cta	tgt	cac	tca	gtg	ccc	aag	gaa	gtg	gtc	cag	ttg	ggt		721
Leu	Leu	Glu	Leu	Cys	His	Ser	Val	Pro	Lys	Glu	Val	Val	Gln	Leu	Gly		
			160					165					170				
gag	gcc	aaa	gat	ggc	agc	aac	ctc	tgc	ttc	agc	aaa	tgattgtgta					767
Glu	Ala	Lys	Asp	Gly	Ser	Asn	Leu	Cys	Phe	Ser	Lys						
		175				180											
accctggggc	acttggtccc	ctctggacct	gattcaccca	tttggaagtt	tgtagcccta												827
gctgatactc	aatggactag	gcctcctcac	ttgtcaatag	tggttccagg	ctgggcgcag												887
tggctcatgc	ctgtgggtccc	ggcacttcgg	gaggccagat	ggggtggctc	acctgagggtc												947
aggagttcga	gaccatcctg	gccaacatgg	tgaaacccca	tctccactaa	aatgcaaaaa												1007
attagccagg	tgtggtggcg	ggcacctgta	gtctcagcta	ctcgggagga	tgaggcagga												1067
aaatcgcttg	aaccagggag	gtggaggttg	cagttgagct	gagatcgtgc	cattgcactc												1127
cagcctgggc	aacgagagca	aaactccatc	tcaaaaaaaa	aaaaaaa													1174

<210> 70  
 <211> 1285  
 <212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 2..1051

<221> sig\_peptide

<222> 2..205

<223> Von Heijne matrix

score 7.3

seq LRLILSPLPGAQP/QQ

<221> polyA\_signal

<222> 1248..1253

<221> polyA\_site

<222> 1272..1285

<400> 70

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Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu	
-65 -60 -55	
gcc tgc agg gcg ctg gtg ttc ggc ggc tgc gtg gag aag tcc tct gtc	97
Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val	
-50 -45 -40	
agc cgc aac cct gag gtg ccc ttt gag agc agt gcc tac cgc atc tca	145
Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser	
-35 -30 -25	
gct tca gcc cgc ggc aag gag ctg cgc ctg ata ctg agc cct ctg cct	193
Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro	
-20 -15 -10 -5	
ggg gcc cag ccc caa cag gag cca ctg gcc ctg gtc ttc cgc ttc ggc	241
Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly	
1 5 10	
atg tcc ggc tct ttt cag ctg gtg ccc cgc gag gag ctg cca cgc cat	289
Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His	
15 20 25	
gcc cac ctg cgc ttt tac acg gcc ccg cct ggc ccc cgg ctc gcc cta	337
Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu	
30 35 40	
tgt ttc gtg gac atc cgc cgg ttc ggc cgc tgg gac ctt ggg gga aag	385
Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys	
45 50 55 60	
tgg cag ccg ggc cgc ggg ccc tgt gtc ttg cag gag tac cag cag ttc	433
Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe	
65 70 75	
agg ctg aag atc ccc ccc ttt gag aag gcc cgc tcg gtc ctg gag gcc	481
Arg Leu Lys Ile Pro Pro Phe Glu Lys Ala Arg Ser Val Leu Glu Ala	
80 85 90	
ctg cag cag cac agg ccg agc ccg gag ctg acc ctg agc cag aag ata	529
Leu Gln Gln His Arg Pro Ser Pro Glu Leu Thr Leu Ser Gln Lys Ile	
95 100 105	
agg acc aag ctg cag aat cca gac ctg ctg gag cta tgt cac tca gtg	577
Arg Thr Lys Leu Gln Asn Pro Asp Leu Leu Glu Leu Cys His Ser Val	
110 115 120	
ccc aag gaa gtg gac cag ttg ggg ggc agg ggc tac ggg tca gag agc	625
Pro Lys Glu Val Asp Gln Leu Gly Gly Arg Gly Tyr Gly Ser Glu Ser	

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125          130          135          140
ggg gag gag gac ttt gct gcc ttt cga gcc tgg ctg cgc tgc tat ggc      673
Gly Glu Glu Asp Phe Ala Ala Phe Arg Ala Trp Leu Arg Cys Tyr Gly
          145          150          155
atg cca ggc atg agc tcc ctg cag gac cgg cat ggc cgt acc atc tgg      721
Met Pro Gly Met Ser Ser Leu Gln Asp Arg His Gly Arg Thr Ile Trp
          160          165          170
ttc cag ggc gat cct gga ccg ttg gca ccc aaa ggc cgc aag tcc cgc      769
Phe Gln Gly Asp Pro Gly Pro Leu Ala Pro Lys Gly Arg Lys Ser Arg
          175          180          185
aaa aag aaa tcc aag gcc aca cag ctg agt cct gag gac aga gtg gag      817
Lys Lys Lys Ser Lys Ala Thr Gln Leu Ser Pro Glu Asp Arg Val Glu
          190          195          200
gac gct ttg cct cca agc aag gcc cct tcc aag aca cga agg gca aag      865
Asp Ala Leu Pro Pro Ser Lys Ala Pro Ser Lys Thr Arg Arg Ala Lys
205          210          215          220
aga gac ctt cct aag agg act gca acc cag cgg cct gag ggg acc agc      913
Arg Asp Leu Pro Lys Arg Thr Ala Thr Gln Arg Pro Glu Gly Thr Ser
          225          230          235
ctc cag cag gac cca gaa gct ccc aca gtg ccc aag aag ggg agg agg      961
Leu Gln Gln Asp Pro Glu Ala Pro Thr Val Pro Lys Lys Gly Arg Arg
          240          245          250
aag ggc cga cag gca gcc tct ggc cac tgc aga ccc cgg aag gtc aag      1009
Lys Gly Arg Gln Ala Ala Ser Gly His Cys Arg Pro Arg Lys Val Lys
          255          260          265
gct gac atc cca tcc ttg gaa cca gag ggg acc tca gcc tct      1051
Ala Asp Ile Pro Ser Leu Glu Pro Glu Gly Thr Ser Ala Ser
          270          275          280
tagcaggagg ctctccttgc ttgcactcac cctttcttat tgtcttgccc tgcattctggg      1111
gggtctgaatt tttgggagca ggcaatatct gaaggtgcaa acaggcccta cggctgttcc      1171
ctgcacaact ctcatggttt taattgtacc ccatcttcca catctttaaa gctcatgtga      1231
aaaatgctgc atttttaata aactgatata tttgaactcc aaaaaaaaaa aaaa      1285

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<210> 71  
<211> 1398  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 2..1171  
<221> sig\_peptide  
<222> 2..205  
<223> Von Heijne matrix  
score 7.3  
seq LRLILSPLPGAQP/QQ

<221> polyA\_signal  
<222> 1368..1373  
<221> polyA\_site  
<222> 1386..1398

<400> 71

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Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu	
-65 -60 -55	
gcc tgc agg gcg ctg gtg ttc ggc ggc tgc gtg gag aag tcc tct gtc	97
Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val	
-50 -45 -40	
agc cgc aac cct gag gtg ccc ttt gag agc agt gcc tac cgc atc tca	145
Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser	
-35 -30 -25	
gct tca gcc cgc ggc aag gag ctg cgc ctg ata ctg agc cct ctg cct	193
Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro	
-20 -15 -10 -5	
ggg gcc cag ccc caa cag gag cca ctg gcc ctg gtc ttc cgc ttc ggc	241
Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly	
1 5 10	
atg tcc ggc tct ttt cag ctg gtg ccc cgc gag gag ctg cca cgc cat	289
Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His	
15 20 25	
gcc cac ctg cgc ttt tac acg gcc ccg cct ggc ccc cgg ctg gcc cta	337
Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu	
30 35 40	
tgt ttc gtg gac atc cgc cgg ttc ggc cgc tgg gac ctt ggg gga aag	385
Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys	
45 50 55 60	
ttg cag ccg ggc cgc ggg ccc tgt gtc ttg cag gag tac cag cag ttc	433
Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe	
65 70 75	
agg gag aat gtg cta cga aac cta gcg gat aag gcc ttt gac cgg ccc	481
Arg Glu Asn Val Leu Arg Asn Leu Ala Asp Lys Ala Phe Asp Arg Pro	
80 85 90	
atc tgc gag gcc ctg ctg gac cag agg ttc ttc aat ggc att ggc aac	529
Ile Cys Glu Ala Leu Leu Asp Gln Arg Phe Phe Asn Gly Ile Gly Asn	
95 100 105	
tat ctg cgg gca gag atc ctg tac cgg ctg aag atc ccc ccc ttt gag	577
Tyr Leu Arg Ala Glu Ile Leu Tyr Arg Leu Lys Ile Pro Pro Phe Glu	
110 115 120	
aag gcc cgc tcg gtc ctg gag gcc ctg cag cag cac agg ccg agc ccg	625
Lys Ala Arg Ser Val Leu Glu Ala Leu Gln Gln His Arg Pro Ser Pro	
125 130 135 140	
gag ctg acc ctg agc cag aag ata agg acc aag ctg cag aat cca gac	673
Glu Leu Thr Leu Ser Gln Lys Ile Arg Thr Lys Leu Gln Asn Pro Asp	
145 150 155	
ctg ctg gag cta tgt cac tca gtg ccc aag gaa gtg gtc cag ttg ggg	721
Leu Leu Glu Leu Cys His Ser Val Pro Lys Glu Val Val Gln Leu Gly	
160 165 170	
ggc aga ggc tac ggg tca gag agc ggg gag gag gac ttt gct gcc ttt	769
Gly Arg Gly Tyr Gly Ser Glu Ser Gly Glu Glu Asp Phe Ala Ala Phe	
175 180 185	
cga gcc tgg ctg cgc tgc tat ggc atg cca ggc atg agc tcc ctg cag	817
Arg Ala Trp Leu Arg Cys Tyr Gly Met Pro Gly Met Ser Ser Leu Gln	
190 195 200	
gac cgg cat ggc cgt acc atc tgg ttc cag ggg gat cct gga ccg ttg	865
Asp Arg His Gly Arg Thr Ile Trp Phe Gln Gly Asp Pro Gly Pro Leu	
205 210 215 220	
gca ccc aaa ggg cgc aag tcc cgc aaa aag aaa tcc aag gcc aca cag	913
Ala Pro Lys Gly Arg Lys Ser Arg Lys Lys Lys Ser Lys Ala Thr Gln	
225 230 235	

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<210> 72
<211> 821
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 42..611

<221> sig_peptide
<222> 42..287
<223> Von Heijne matrix
      score 4.4
      seq NLPHLQVVGLTWG/HI

<221> polyA_signal
<222> 787..792

<221> polyA_site
<222> 808..821

<400> 72
ccggttgccag ttctgcgcgt gtcctgcgtc tccagtatgg a atg tat gtt tgg ccc          56
                                     Met Tyr Val Trp Pro
                                     -80
tgt gct gtg gtc ctg gcc cag tac ctt tgg ttt cac aga aga tct ctg          104
Cys Ala Val Val Leu Ala Gln Tyr Leu Trp Phe His Arg Arg Ser Leu
      -75                                -70                                -65
cca ggc aag gcc atc tta gag att gga gca gga gtg agc ctt cca gga          152
Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala Gly Val Ser Leu Pro Gly
      -60                                -55                                -50
att ttg act gcc aaa tgt ggt gca gaa gta ata ctg tca gac aac tca          200

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^220>
^221> CDS
^222> 42..611
^223> sig_peptide
^224> 42..287
^225> Von Heijne matrix
      score 4.4
      seq NLPHLOVVGLTWG/HI

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<221> polyA_signal
<222> 787..792
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<221> polyA_site
<222> 808..821
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[illegible]

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Ile Leu Thr Ala Lys Cys Gly Ala Glu Val Ile Leu Ser Asp Ser Ser
-45          -40          -35          -30
gaa ctg cct cac tgt ctg gaa gtc tgt cgg caa agc tgc caa atg aat      248
Glu Leu Pro His Cys Leu Glu Val Cys Arg Gln Ser Cys Gln Met Asn
          -25          -20          -15
aac ctg cca cat ctg cag gtg gta gga cta aca tgg ggt cat ata tct      296
Asn Leu Pro His Leu Gln Val Val Gly Leu Thr Trp Gly His Ile Ser
          -10          -5          1
tgg gat ctt ctg gct cta cca cca caa gat att atc ctt gca tct gat      344
Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp Ile Ile Leu Ala Ser Asp
          5          10          15
gtg ttc ttt gaa cca gaa gat ttt gaa gac att ttg gct aca ata tat      392
Val Phe Phe Glu Pro Glu Asp Phe Glu Asp Ile Leu Ala Thr Ile Tyr
          20          25          30          35
ttt ttg atg cac aag aat ccc aag gtc caa ttg tgg tct act tat caa      440
Phe Leu Met His Lys Asn Pro Lys Val Gln Leu Trp Ser Thr Tyr Gln
          40          45          50
gtt agg agt gct gac tgg tca ctt gaa gct tta ctc tac aaa tgg gat      488
Val Arg Ser Ala Asp Trp Ser Leu Glu Ala Leu Leu Tyr Lys Trp Asp
          55          60          65
atg aaa tgt gtc cac att cct ctt gag tct ttt gat gca gac aaa gaa      536
Met Lys Cys Val His Ile Pro Leu Glu Ser Phe Asp Ala Asp Lys Glu
          70          75          80
gat ata gca gaa tct acc ctt cca gga aga cat aca gtt gaa atg ctg      584
Asp Ile Ala Glu Ser Thr Leu Pro Gly Arg His Thr Val Glu Met Leu
          85          90          95
gtc att tcc ttt gca aag gac agt ctc tgaattatac ctacaacctg      631
Val Ile Ser Phe Ala Lys Asp Ser Leu
          100          105
ttctgggaca gtatcaatac tgatgagcaa cctggcacac aaactatgag cagaccactt      691
cagcttgaga atgcagtggg tctgaagatg gtcaagtctg tctgccttag attttgatgt      751
cacctagaca acacttaaac tcatatgaaa caaaaattaa aatacgtatt acaagtaaaa      811
aaaaaaaaaa      821

<210> 73
<211> 916
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 62..916

<221> sig_peptide
<222> 62..757
<223> Von Heijne matrix
      score 4.2
      seq LVTPAALRPLVLG/GN

<221> polyA_site
<222> 904..916

<400> 73
cctgaatgac ttgaatgttt ccccgctga gctaacagtc catgtgggtg attcagctct      60
g atg gga tgt gtt ttc cag agc aca gaa gac aaa cgt ata ttc aag ata      109

```



Met	Gly	Cys	Val	Phe	Gln	Ser	Thr	Glu	Asp	Lys	Arg	Ile	Phe	Lys	Ile			
-230																-225	-220	
gac	tgg	act	ctg	tca	cca	gga	gag	cac	gcc	aag	gac	gaa	tat	gtg	cta	157		
Asp	Trp	Thr	Leu	Ser	Pro	Gly	Glu	His	Ala	Lys	Asp	Glu	Tyr	Val	Leu			
-215																-210	-205	
tac	tat	tac	tcc	aat	ctc	agt	gtg	cct	att	ggg	cgc	ttc	cag	aac	cgc	205		
Tyr	Tyr	Tyr	Ser	Asn	Leu	Ser	Val	Pro	Ile	Gly	Arg	Phe	Gln	Asn	Arg			
-200																-195	-190	-185
gta	cac	ttg	atg	ggg	gac	aac	tta	tgc	aat	gat	ggc	tct	ctc	ctg	ctc	253		
Val	His	Leu	Met	Gly	Asp	Asn	Leu	Cys	Asn	Asp	Gly	Ser	Leu	Leu	Leu			
-180																-175	-170	
caa	gat	gtg	caa	gag	gct	gac	cag	gga	acc	tat	atc	tgt	gaa	atc	cgc	301		
Gln	Asp	Val	Gln	Glu	Ala	Asp	Gln	Gly	Thr	Tyr	Ile	Cys	Glu	Ile	Arg			
-165																-160	-155	
ctc	aaa	ggg	gag	agc	cag	gtg	ttc	aag	aag	gcg	gtg	gta	ctg	cat	gtg	349		
Leu	Lys	Gly	Glu	Ser	Gln	Val	Phe	Lys	Lys	Ala	Val	Val	Leu	His	Val			
-150																-145	-140	
ctt	cca	gag	gag	ccc	aaa	gag	ctc	atg	gtc	cat	gtg	ggg	gga	ttg	att	397		
Leu	Pro	Glu	Glu	Pro	Lys	Glu	Leu	Met	Val	His	Val	Gly	Gly	Leu	Ile			
-135																-130	-125	
cag	atg	gga	tgt	gtt	ttc	cag	agc	aca	gaa	gtg	aaa	cac	gtg	acc	aag	445		
Gln	Met	Gly	Cys	Val	Phe	Gln	Ser	Thr	Glu	Val	Lys	His	Val	Thr	Lys			
-120																-115	-110	-105
gta	gaa	tgg	ata	ttt	tca	gga	cgg	cgc	gca	aag	gag	gag	att	gta	ttt	493		
Val	Glu	Trp	Ile	Phe	Ser	Gly	Arg	Arg	Ala	Lys	Glu	Glu	Ile	Val	Phe			
-100																-95	-90	
cgt	tac	tac	cac	aaa	ctc	agg	atg	tct	gcg	gag	tac	tcc	cag	agc	tgg	541		
Arg	Tyr	Tyr	His	Lys	Leu	Arg	Met	Ser	Ala	Glu	Tyr	Ser	Gln	Ser	Trp			
-85																-80	-75	
ggc	cac	ttc	cag	aat	cgt	gtg	aac	ctg	gtg	ggg	gac	att	ttc	cgc	aat	589		
Gly	His	Phe	Gln	Asn	Arg	Val	Asn	Leu	Val	Gly	Asp	Ile	Phe	Arg	Asn			
-70																-65	-60	
gac	ggg	tcc	atc	atg	ctt	caa	gga	gtg	agg	gag	tca	gat	gga	gga	aac	637		
Asp	Gly	Ser	Ile	Met	Leu	Gln	Gly	Val	Arg	Glu	Ser	Asp	Gly	Gly	Asn			
-55																-50	-45	
tac	acc	tgc	agt	atc	cac	cta	ggg	aac	ctg	gtg	ttc	aag	aaa	acc	att	685		
Tyr	Thr	Cys	Ser	Ile	His	Leu	Gly	Asn	Leu	Val	Phe	Lys	Lys	Thr	Ile			
-40																-35	-30	-25
gtg	ctg	cat	gtc	agc	ccg	gaa	gag	cct	cga	aca	ctg	gtg	acc	ccg	gca	733		
Val	Leu	His	Val	Ser	Pro	Glu	Glu	Pro	Arg	Thr	Leu	Val	Thr	Pro	Ala			
-20																-15	-10	
gcc	ctg	agg	cct	ctg	gtc	ttg	ggg	ggg	aat	cag	ttg	gtg	atc	att	gtg	781		
Ala	Leu	Arg	Pro	Leu	Val	Leu	Gly	Gly	Asn	Gln	Leu	Val	Ile	Ile	Val			
-5																1	5	
gga	att	gtc	tgt	gcc	aca	atc	ctg	ctg	ctc	cct	gtc	ctg	ata	ttg	atc	829		
Gly	Ile	Val	Cys	Ala	Thr	Ile	Leu	Leu	Leu	Pro	Val	Leu	Ile	Leu	Ile			
10																15	20	
gtg	aag	aag	acc	tgt	gga	aat	aag	agt	tca	gtg	aat	tct	aca	gtc	ttg	877		
Val	Lys	Lys	Thr	Cys	Gly	Asn	Lys	Ser	Ser	Val	Asn	Ser	Thr	Val	Leu			
25																30	35	40
gtg	aag	aac	acg	aag	aag	act	aat	cca	aaa	aaa	aaa	aaa	aaa	aaa	aaa	916		
Val	Lys	Asn	Thr	Lys	Lys	Thr	Asn	Pro	Lys	Lys	Lys	Lys	Lys	Lys	Lys			
45																50		

<211> 1153  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 62..520

<221> polyA\_signal  
<222> 1124..1129

<221> polyA\_site  
<222> 1141..1153

<400> 74

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g atg gga tgt gtt ttc cag agc aca gta gac aaa tgt ata ttc aag ata      109
  Met Gly Cys Val Phe Gln Ser Thr Val Asp Lys Cys Ile Phe Lys Ile
    1             5             10             15
gac tgg act ctg tca cca gga gag cac gcc aag gac gaa tat gtg cta      157
Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu
    20             25             30
tac tat tac tcc aat ctc agt gtg cct att ggg cgc ttc cag aac cgc      205
Tyr Tyr Tyr Ser Asn Leu Ser Val Pro Ile Gly Arg Phe Gln Asn Arg
    35             40             45
gta cac ttg atg ggg gac atc tta tgc aat gat ggc tct ctc ctg ctc      253
Val His Leu Met Gly Asp Ile Leu Cys Asn Asp Gly Ser Leu Leu Leu
    50             55             60
caa gat gtg caa gag gct gac cag gga acc tat atc tgt gaa atc cgc      301
Gln Asp Val Gln Glu Ala Asp Gln Gly Thr Tyr Ile Cys Glu Ile Arg
    65             70             75             80
ctc aaa ggg gag agc cag gtg ttc aag aag gcg gtg gta ctg cat gtg      349
Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val
    85             90             95
ctt cca gag gag ccc aaa gag ctc atg gtc cat gtg ggt gga ttg att      397
Leu Pro Glu Glu Pro Lys Glu Leu Met Val His Val Gly Leu Ile
    100            105            110
cag atg gga tgt gtt ttc cag agc aca gaa gtg aaa cac gtg acc aag      445
Gln Met Gly Cys Val Phe Gln Ser Thr Glu Val Lys His Val Thr Lys
    115            120            125
gta gaa tgg ata ttt tca gga cgg cgc gca aag gta aca agg agg aaa      493
Val Glu Trp Ile Phe Ser Gly Arg Arg Ala Lys Val Thr Arg Arg Lys
    130            135            140
cat cac tgt gtt aga gaa ggc tct ggc tgatggtatc aggacaaagg      540
His His Cys Val Arg Glu Gly Ser Gly
    145            150
tagaatcagg cacatgagga ggtgttgcaa gaggctgggc tttggtgctt atcagaactg      600
gaccttctcc tagcaatttc agctttcttg tgggaaaggt aactccaatg aagaacaaga      660
acaagaagat gatgatgatg cttaactttt tggatgccga tatgagattg tacatgtaaa      720
gcattttgta taagacttgg ccctgcatt ttagtttctt tctttctccc ttttctctcg      780
tatagatgcc atgggagaaat gagggagatg atttttgtgg ccagccaag aaagcaatgg      840
gctagacatt aaaatgatta cacttttatt cttactgggg ttagttctgt gagttttcat      900
ctgtgccccca ttgccccatt tatgtgatgg agggaatttt catgggtact tcacgtgttg      960
ggattgattg atcctggggg ccagggtgaa ggtattttta cgggacctct ataaagcagg      1020
aagaagcaag tttattcttt agaccagtag ctctcaacca tgatgtggtc atatatttat      1080
gggtcaacat gtgttggtgg gatatcccaa gtaacttggt attaataaaa gttaagttgc      1140
aaaaaaaaaa aaa
1153
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<210> 75  
 <211> 1517  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 21..167

<400> 75

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ctctgaaatg cttgtctttt atg ctg gna ggt gac cat agg gct ctg ctt tta      53
                Met Leu Xaa Gly Asp His Arg Ala Leu Leu Leu
                1          5          10
aag ata tgg ctg ctt caa agg cca gag tca cag gaa gga ctt ctt cca      101
Lys Ile Trp Leu Leu Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro
                15          20          25
ggg aga tta gtg gtg atg gag agg aga gtt aaa atg acc tca tgt cct      149
Gly Arg Leu Val Val Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro
                30          35          40
tct tgt cca cgg ttt tgt tgagttttca ctctttctaata gcaaggggtct      197
Ser Cys Pro Arg Phe Cys
                45
cacactgtga accacttagg atgtgatcac tttcaggtgg ccaggaatgt tgaatgtctt      257
tggtctcagtt catttaaaaa agatatctat ttgaaagtgc tcagagttgt acatatgttt      317
cacagtacag gatctgtaca taaaagtttc tttcctaaac cattcaccaa gagccaatat      377
ctaggcattt tcttggtagc acaaattttc ttattgctta gaaaattgtc ctccttggtta      437
tttctgtttg taagacttaa gtgagttagg tctttaagga aagcaacgct cctctgaaat      497
gcttgtcttt tatgctggga ggtgaccata gggctctgct tttaaagata tggctgcttc      557
aaaggccaga gtcacaggaa ggacttcttc cagggagatt agtgggtgat gagaggagag      617
ttaaaatgac ctcatgtcct tcttgtccac ggttttggtg agttttcact cttctaattgc      677
aagggtctca cactgtgaac cacttaggat gtgatcactt tcaggtggcc aggaatgttg      737
aatgtctttg gctcagttca tttaaaaaag atatctattt gaaagttctc agagttgtac      797
tatatgtttca cagtacagga tctgtacata aaagtttctt tcctaaacca ttcaccaaga      857
gccaatatct aggcattttc ttggtagcac aaattttctt attgcttaga aaattgtcct      917
ccttgtttatt tctgtttgta agacttaagt gagttaggtc ttttaaggaaa gcaacgctcc      977
tctgaaatgc ttgtcttttna tgctgggagg tgaccatagg gctctgcttt taaagatatg      1037
gctgcttcaa aggccagagt cacaggaagg acttcttcca gggagattag tggatgatga      1097
gaggagagtt aaaatgacct catgtccttc ttgtccacgg ttttgttgag ttttcaactc      1157
tctaatacaa gggcttcaca ctgtgaacca cttaggatgt gatcactttc aggtggccag      1217
gaatgttgaa tgtctttggc tcagttcatt taaaaaagat atctatttga aagttctcag      1277
agttgtacat atgtttcaca gtacaggatc tgtacataaa agtttctttc ctaaaccatt      1337
caccaagagc caatatctag gcattttctt ggtagcacia attttcttat tgcttagaaa      1397
attgtcctcc ttgttatctc tgtttgtaag acttaagtga gttaggtctt taaggaaagc      1457
aacgctcctc tgaaatgctt gtcttttatg ctgggaggtg accatagggc tctgctttta      1517

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<210> 76  
 <211> 526  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS

<222> 22..318

<221> sig\_peptide

<222> 22..93

<223> Von Heijne matrix

score 4.6

seq FFIFCSLNTLLLG/GV

<221> polyA\_signal

<222> 497..502

<221> polyA\_site

<222> 516..526

<400> 76

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ctgcctgctg cttgctgcac c atg aag tct gcc aag ctg gga ttt ctt cta      51
                               Met Lys Ser Ala Lys Leu Gly Phe Leu Leu
                               -20                               -15
aga ttc ttc atc ttc tgc tca ttg aat acc ctg tta ttg ggt ggt gtt      99
Arg Phe Phe Ile Phe Cys Ser Leu Asn Thr Leu Leu Leu Gly Gly Val
                               -10                               -5                               1
aat aaa att gcg gag aag ata tgt gga gac ctc aaa gat ccc tgc aaa      147
Asn Lys Ile Ala Glu Lys Ile Cys Gly Asp Leu Lys Asp Pro Cys Lys
                               5                               10                               15
ttg gac atg aat ttt gga agc tgc tat gaa gtt cac ttt aga tat ttc      195
Leu Asp Met Asn Phe Gly Ser Cys Tyr Glu Val His Phe Arg Tyr Phe
                               20                               25                               30
tac aac aga acc tcc aaa aga tgt gaa act ttt gtc ttc tcc ggc tgt      243
Tyr Asn Arg Thr Ser Lys Arg Cys Glu Thr Phe Val Phe Ser Gly Cys
                               35                               40                               45                               50
aat ggc aac ctt aac aac ttc aag ctt aaa ata gaa cgt gaa gta gcc      291
Asn Gly Asn Leu Asn Asn Phe Lys Leu Lys Ile Glu Arg Glu Val Ala
                               55                               60                               65
tgt gtt gca aaa tac aaa cca ccg agg tgagaggatg tgaactcatg      338
Cys Val Ala Lys Tyr Lys Pro Pro Arg
                               70                               75
aagttgtctg ctgcaccatc cgaaataaag acacaagaaa attcagactg attttgaaat      398
ctttgtaata tttccataat gctttaagct tccatatgtt tgctattttc ctgaccctag      458
ttttgtcttt cctggaaatt aactgtatga tcattagaat gaaagagtct ttctgtcaaa      518
aaaaaaaaa                                                                526
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<210> 77

<211> 352

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 8..292

<221> sig\_peptide

<222> 8..118

<223> Von Heijne matrix

score 5.6

seq WLLLDALLRLGDT/KK

<221> polyA\_signal  
<222> 317..322

<221> polyA\_site  
<222> 339..352

<400> 77

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ctgagat atg gca agt ccc gct gta aac agg tgg aaa agg cca agg ttg      49
      Met Ala Ser Pro Ala Val Asn Arg Trp Lys Arg Pro Arg Leu
            -35                      -30                      -25
aag ccg gtg tgg cca cgg cgc ttg gaa tcc tgg ttg ttg ctg gat gct      97
Lys Pro Val Trp Pro Arg Arg Leu Glu Ser Trp Leu Leu Leu Asp Ala
            -20                      -15                      -10
ctt ttg cga tta gga gat acc aaa aaa aag cga cag cct gaa gca gcc     145
Leu Leu Arg Leu Gly Asp Thr Lys Lys Lys Arg Gln Pro Glu Ala Ala
            -5                      1                      5
aca aaa tcc tgt gtt aga agc agc tgt ggg ggt ccc agt gga gat ggg     193
Thr Lys Ser Cys Val Arg Ser Ser Cys Gly Gly Pro Ser Gly Asp Gly
10                      15                      20                      25
cct ccc cca tgc ctc cag cag cct gac cct cgt gcc ctg tct cag gcg     241
Pro Pro Pro Cys Leu Gln Gln Pro Asp Pro Arg Ala Leu Ser Gln Ala
            30                      35                      40
ttc tct aga tcc ttt cct ctg ttt ccc tct ctc gct ggc aaa agt atg     289
Phe Ser Arg Ser Phe Pro Leu Phe Pro Ser Leu Ala Gly Lys Ser Met
            45                      50                      55
atc taattgaaac aagactgaag gatcaataaa cagccatctg ccccttcaaa     342
Ile
aaaaaaaaaaaaa                                           352

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<210> 78

<211> 542

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 16..378

<221> sig\_peptide

<222> 16..84

<223> Von Heijne matrix

score 9.8

seq FLLFFFLFLLTRG/SL

<221> polyA\_signal

<222> 502..507

<221> polyA\_site

<222> 522..542

<400> 78

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cacgacctgt gggcc atg atg cta ccc caa tgg ctg ctg ctg ctg ttc ctt      51
      Met Met Leu Pro Gln Trp Leu Leu Leu Leu Phe Leu
            -20                      -15

```

```

ctc ttc ttc ttt ctc ttc ctc ctc acc agg ggc tca ctt tct cca aca      99
Leu Phe Phe Phe Leu Phe Leu Leu Thr Arg Gly Ser Leu Ser Pro Thr
-10 -5 1 5
aaa tat aac ctt ttg gag ctc aag gag tct tgc atc cgg aac cag gac      147
Lys Tyr Asn Leu Leu Glu Leu Lys Glu Ser Cys Ile Arg Asn Gln Asp
10 15 20
tgc gag act ggc tgc tgc caa cgt gct cca gac aat tgc gag tgc cac      195
Cys Glu Thr Gly Cys Cys Gln Arg Ala Pro Asp Asn Cys Glu Ser His
25 30 35
tgc gcg gag aag ggg tcc gag ggc agt ctg tgt caa acg cag gtg ttc      243
Cys Ala Glu Lys Gly Ser Glu Gly Ser Leu Cys Gln Thr Gln Val Phe
40 45 50
ttt ggc caa tat aga gcg tgt ccc tgc ctg cgg aac ctg act tgt ata      291
Phe Gly Gln Tyr Arg Ala Cys Pro Cys Leu Arg Asn Leu Thr Cys Ile
55 60 65
tat tca aag aat gag aaa tgg ctt agc atc gcc tat ggc cgt tgt cag      339
Tyr Ser Lys Asn Glu Lys Trp Leu Ser Ile Ala Tyr Gly Arg Cys Gln
70 75 80 85
aaa att gga agg cag aag ttg gct aag aaa atg ttc ttc tagtgctccc      388
Lys Ile Gly Arg Gln Lys Leu Ala Lys Lys Met Phe Phe
90 95
tcctttcttgc tgctctctcc tctccacct gctctctctcc ctaccagag ctctgtgttc      448
accctgttcc ccagagcctc caccatgagt ggagggaagt ggggagtgat tgaaataaag      508
agcttttttca atgaaaaaaaa aaaaaaaaaa aaaa      542

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<210> 79

<211> 233

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 57..233

<400> 79

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gcacaaaccaa aaccagcacc gatcccgaca tagatcagtg acgtcttttt cttcag atg      59
Met
1
atc cta tgt ttc ctt ctt cct cat cat cgt ctt cag gaa gcc aga cag      107
Ile Leu Cys Phe Leu Leu Pro His His Arg Leu Gln Glu Ala Arg Gln
5 10 15
att caa gta ttg aag atg ctg cca agg gaa aaa tta aga aga aga gaa      155
Ile Gln Val Leu Lys Met Leu Pro Arg Glu Lys Leu Arg Arg Arg Glu
20 25 30
gag aga aaa caa ata aat ggg aaa aaa gaa agg aca aaa tat gaa aca      203
Glu Arg Lys Gln Ile Asn Gly Lys Lys Glu Arg Thr Lys Tyr Glu Thr
35 40 45
cca aga aaa aga gaa gga aaa aaa aaa aaa
Pro Arg Lys Arg Glu Gly Lys Lys Lys Lys      233
50 55

```

<210> 80

<211> 660

<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 83..340

<221> sig\_peptide  
<222> 83..124  
<223> Von Heijne matrix  
score 7.5  
seq VALNLILVPCCAA/WC

<221> polyA\_signal  
<222> 573..578

<221> polyA\_site  
<222> 607..660

<400> 80  
gaatttgtaa aacttctgct cgtttacact gcacattgaa tacaggtaac taattggaag 60  
gagagggggag atcactcttt tg atg gtg gcc ctg aac ctc att ctg gtt ccc 112  
Met Val Ala Leu Asn Leu Ile Leu Val Pro  
-10 -5  
gtgc tgc gct gct tgg tgt gac cca cgg agg atc cac tcc cag gat gac 160  
Cys Cys Ala Ala Trp Cys Asp Pro Arg Arg Ile His Ser Gln Asp Asp  
1 5 10  
gtg ccc cgt agc tct gct gct gat act ggg tct gcg atg cag cgg cgt 208  
Val Pro Arg Ser Ser Ala Ala Asp Thr Gly Ser Ala Met Gln Arg Arg  
15 20 25  
gag gcc tgg gct ggt tgg aga agg tca caa ccc ttc tct gtt ggt ctg 256  
Glu Ala Trp Ala Gly Trp Arg Arg Ser Gln Pro Phe Ser Val Gly Leu  
30 35 40  
tct tct gct gaa aga ctc gag aac caa cca ggg aag ctg tcc tgg agg 304  
Pro Ser Ala Glu Arg Leu Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg  
45 50 55 60  
ccc ctg gtc gga gag gga tat aga atc tgt gac ctc tgacaactgt 350  
Ser Leu Val Gly Glu Gly Tyr Arg Ile Cys Asp Leu  
65 70  
gaagccaccc tgggctacag aaaccacagt cttcccagca attattacaa ttcttgaatt 410  
ccttgggggat tttttactgc cttttcaaag cacttaagtg ttagatctaa cgtgttccag 470  
tgtctgtctg aggtgactta aaaaatcaga acaaaacttc tattatccag agtcatggga 530  
gagtacaccc ttccaggaa taatgttttg ggaaacactg aaatgaaatc ttcccagtat 590  
tataaattgt gtatttataaa aaagaaactt ttctgaatgc ctacctggcg gtgtatacca 650  
ggcagtgtgc 660

<210> 81  
<211> 605  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 47..541

<221> sig\_peptide  
 <222> 47..220  
 <223> Von Heijne matrix  
 score 5.4  
 seq QLLDSVLWLGLG/LT

<221> polyA\_site  
 <222> 597..605

<400> 81

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aaagtgggag gagcactagg tcttcccgtc acctccacct ctctcc atg acc cgg      55
                                     Met Thr Arg
ctc tgc tta ccc aga ccc gaa gca cgt gag gat ccg atc cca gtt cct      103
Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro Ile Pro Val Pro
-55                               -50                               -45
cca agg ggc ctg ggt gct ggg gag ggg tca ggt agt cca gtg cgt cca      151
Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser Pro Val Arg Pro
                               -35                               -30                               -25
cct gta tcc acc tgg ggc cct agc tgg gcc cag ctc ctg gac agt gtc      199
Pro Val Ser Thr Trp Gly Pro Ser Trp Ala Gln Leu Leu Asp Ser Val
                               -20                               -15                               -10
cta tgg ctg ggg gca cta gga ctg aca atc cag gca gtc ttt tcc acc      247
Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln Ala Val Phe Ser Thr
-5                               1                               5
act ggc cca gcc ctg ctg ctg ctt ctg gtc agc ttc ctc acc ttt gac      295
Thr Gly Pro Ala Leu Leu Leu Leu Leu Val Ser Phe Leu Thr Phe Asp
10                               15                               20                               25
ctg ctc cat agg ccc gca ggt cac act ctg cca cag cgc aaa ctt ctc      343
Leu Leu His Arg Pro Ala Gly His Thr Leu Pro Gln Arg Lys Leu Leu
                               30                               35                               40
acc agg ggc cag agt cag ggg gcc ggt gaa ggt cct gga cag cag gag      391
Thr Arg Gly Gln Ser Gln Gly Ala Gly Glu Gly Pro Gly Gln Gln Glu
                               45                               50                               55
gct cta ctc ctg caa atg ggt aca gtc tca gga caa ctt agc ctc cag      439
Ala Leu Leu Leu Gln Met Gly Thr Val Ser Gly Gln Leu Ser Leu Gln
                               60                               65                               70
gac gca ctg ctg ctg ctg ctc atg ggg ctg ggc ccg ctc ctg aga gcc      487
Asp Ala Leu Leu Leu Leu Leu Met Gly Leu Gly Pro Leu Leu Arg Ala
75                               80                               85
tgt ggc atg ccc ttg acc ctg ctt ggc ctg gct ttc tgc ctc cat cct      535
Cys Gly Met Pro Leu Thr Leu Leu Gly Leu Ala Phe Cys Leu His Pro
90                               95                               100                               105
tgg gcc tgagagcccc tccccacaac tcagtgtcct tcaaataac aatgaccacc      591
Trp Ala
cttctttcaaa aaaa      605

```

<210> 82  
 <211> 396  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 46..285



<221> sig\_peptide  
 <222> 46..150  
 <223> Von Heijne matrix  
 score 3.6  
 seq LEPGLSSSAACNG/KE

<221> polyA\_signal  
 <222> 364..369

<221> polyA\_site  
 <222> 385..396

<400> 82

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cctctacagg aatcagactc agcctctttt gggttttcagt gaagt atg cct ttt caa      57
                                     Met Pro Phe Gln
                                     -35
ttt gga acc cag cca agg agg ttt cca gtg gaa gga gga gat tct tca      105
Phe Gly Thr Gln Pro Arg Arg Phe Pro Val Glu Gly Gly Asp Ser Ser
  -30                -25                -20
att gag ctg gaa cct ggg ctg agc tcc agt gct gcc tgt aat ggg aag      153
Ile Glu Leu Glu Pro Gly Leu Ser Ser Ser Ala Ala Cys Asn Gly Lys
  -15                -10                -5                1
gag atg tca cca acc agg caa ctc cgg agg tgc cct gga agt cat tgc      201
Glu Met Ser Pro Thr Arg Gln Leu Arg Arg Cys Pro Gly Ser His Cys
          5                10                15
ctg aca ata act gat gtt ccc gtc act gtt tat gca aca acg aga aag      249
Leu Thr Ile Thr Asp Val Pro Val Thr Val Tyr Ala Thr Thr Arg Lys
    20                25                30
cca cct gca caa agc agc aag gaa atg cat cct aaa tagcaccatt      295
Pro Pro Ala Gln Ser Ser Lys Glu Met His Pro Lys
    35                40                45
aagtcttttg tcaaggtctg actaggtcaa gggtaatgga ccagtatcat ctggtgatct      355
ggtaaacaata taaaagtggt ggcaccttca aaaaaaaaaa a      396
```

<210> 83

<211> 432

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 22..240

<221> sig\_peptide

<222> 22..84

<223> Von Heijne matrix  
 score 12  
 seq VLVLCLVLLQLAQG/GY

<221> polyA\_signal

<222> 397..402

<221> polyA\_site

<222> 421..432

<400> 83

```

gctcacgctc tgggtcagagt t atg gca ccc cag act ctg ctg cct gtc ctg      51
                        Met Ala Pro Gln Thr Leu Leu Pro Val Leu
                        -20                                -15
gtt ctc tgt gtg ctg ctg ctg cag gcc cag gga gga tac cgt gac aag      99
Val Leu Cys Val Leu Leu Leu Gln Ala Gln Gly Gly Tyr Arg Asp Lys
-10                    -5                                1                    5
atg agg atg cag aga atc aag gtc tgt gag aag cga ccc agc ata gat      147
Met Arg Met Gln Arg Ile Lys Val Cys Glu Lys Arg Pro Ser Ile Asp
                        10                    15                    20
cta tgc atc cac cac tgt tca tgt ttc caa aag tgt gaa aca aat aag      195
Leu Cys Ile His His Cys Ser Cys Phe Gln Lys Cys Glu Thr Asn Lys
                        25                    30                    35
ata tgc tgt tca gcc ttc tgt ggg aac att tgt atg agc atc cta      240
Ile Cys Cys Ser Ala Phe Cys Gly Asn Ile Cys Met Ser Ile Leu
                        40                    45                    50
tgagtgggag agtgggctgg gatgtgcatc ctgctccctg aacccttcca tccgagactg      300
tgcccacatc cgaagcacaa ggacatcaaa tcatcagcac aagaacatca acaggaatgc      360
caccctcccc agtgtctgaa ctccctgtcc ctgtcaaatg aaccagaaca aatgccccatg      420
aaaaaaaaaa aa                                                         432

```

<210> 84

<211> 420

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 89..382

<221> polyA\_site

<222> 408..420

<400> 84

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gcttgctga ccccatgtc gcctctgtag gtagaagaag tatgtcttcc tggacccct      60
ggctggtgct gtaacaaaga cccatgtg atg ctg ggg gca gag aca gag gag      112
                        Met Leu Gly Ala Glu Thr Glu Glu
                        1                    5
aag ctg ttt gat gcc ccc ttg tcc atc agc aag aga gag cag ctg gaa      160
Lys Leu Phe Asp Ala Pro Leu Ser Ile Ser Lys Arg Glu Gln Leu Glu
                        10                    15                    20
cag cag gtc cca gag aac tac ttc tat gtg cca gac ctg ggc cag gtg      208
Gln Gln Val Pro Glu Asn Tyr Phe Tyr Val Pro Asp Leu Gly Gln Val
                        25                    30                    35                    40
cct gag att gat gtt cca tcc tac ctg cct gac ctg ccc ggc att gcc      256
Pro Glu Ile Asp Val Pro Ser Tyr Leu Pro Asp Leu Pro Gly Ile Ala
                        45                    50                    55
aac gac ctc atg tac att gcc gac ctg ggc ccc ggc att gcc ccc tct      304
Asn Asp Leu Met Tyr Ile Ala Asp Leu Gly Pro Gly Ile Ala Pro Ser
                        60                    65                    70
gcc cct ggc acc att cca gaa ctg ccc acc ttc cac act gag gta gcc      352
Ala Pro Gly Thr Ile Pro Glu Leu Pro Thr Phe His Thr Glu Val Ala
                        75                    80                    85
gag cct ctc aag acc tac aag atg ggg tac taacagcacc accacogccc      402
Glu Pro Leu Lys Thr Tyr Lys Met Gly Tyr

```

90  
ccaccaaaaa aaaaaaaa

95

420

<210> 85  
<211> 501  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 80..415

<221> sig\_peptide  
<222> 80..142  
<223> Von Heijne matrix  
score 5.4  
seq TFCLIFGLGAVWG/LG

<221> polyA\_signal  
<222> 471..476

<221> polyA\_site  
<222> 488..501

<400> 85

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cccgcttgat tccaagaacc tcttcgatat ttatTTTTat ttttaaagag ggagacgatg      60
gactgagctg atccgcacc atg gag tct cgg gtc tta ctg aga aca ttc tgt      112
                Met Glu Ser Arg Val Leu Leu Arg Thr Phe Cys
                -20                      -15
ttg atc ttc ggt ctc gga gca gtt tgg ggg ctt ggt gtg gac cct tcc      160
Leu Ile Phe Gly Leu Gly Ala Val Trp Gly Leu Gly Val Asp Pro Ser
-10                      -5                      1                      5
cta cag att gac gtc tta aca gag tta gaa ctt ggg gag tcc acg acc      208
Leu Gln Ile Asp Val Leu Thr Glu Leu Glu Leu Gly Glu Ser Thr Thr
                10                      15                      20
gga gtg cgt cag gtc ccg ggg ctg cat aat ggg acg aaa gcc ttt ctc      256
Gly Val Arg Gln Val Pro Gly Leu His Asn Gly Thr Lys Ala Phe Leu
                25                      30                      35
ttt caa gat act ccc aga agc ata aaa gca tcc act gct aca gct gaa      304
Phe Gln Asp Thr Pro Arg Ser Ile Lys Ala Ser Thr Ala Thr Ala Glu
                40                      45                      50
cag ttt ttt cag aag ctg aga aat aaa cat gaa ttt act att ttg gtg      352
Gln Phe Phe Gln Lys Leu Arg Asn Lys His Glu Phe Thr Ile Leu Val
55                      60                      65                      70
acc cta aaa cag acc cac tta aat tca gga gtt att ctc tca att cac      400
Thr Leu Lys Gln Thr His Leu Asn Ser Gly Val Ile Leu Ser Ile His
                75                      80                      85
cac ttg gat cac agg taaatgtggg tgctggagtt tcctgtgttt tcattatatg      455
His Leu Asp His Arg
                90
tgggttaaatg aatatattaa agagaagtaa acaaaaaaaaa aaaaaa      501
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<210> 86

<211> 454  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 152..361

<221> sig\_peptide  
<222> 152..283  
<223> Von Heijne matrix  
score 4.7  
seq FLLSLSLITYCFW/DP

<400> 86  
gacatttttac ttttttctgt taacgcttac cctagaaatt agaaatgaca ccacgtattc 60  
ttagcgaagt ccagttttca gcattttgtc cttattggac aatagcaagg atattagaac 120  
gtgttggttc cgcgtgcttc cgtcttgagt t atg tgc tgc tat tgt cgg ata 172  
Met Cys Cys Tyr Cys Arg Ile  
-40  
ttt tgt ctt aga tgt acg tac ttt cct gtt cat tgt ggt atg tgt aat 220  
Phe Cys Leu Arg Cys Thr Tyr Phe Pro Val His Cys Gly Met Cys Asn  
-35 -30 -25  
ttg cgt tac ttt gaa ttt tcc acg ttt tta ctt tct ttg tct ctc atc 268  
Leu Arg Tyr Phe Glu Phe Ser Thr Phe Leu Leu Ser Leu Ser Leu Ile  
-20 -15 -10  
act tac tgc ttt tgg gac ccc ccc cat cgg ggt tca cat tcc ctc tcc 316  
Thr Tyr Cys Phe Trp Asp Pro Pro His Arg Gly Ser His Ser Leu Ser  
-5 1 5 10  
cta gag cac act ccc ttg gat ttc ctc gag tgg ggt ctg ctg cgg 361  
Leu Glu His Thr Pro Leu Asp Phe Leu Glu Trp Gly Leu Leu Arg  
15 20 25  
tgaagctttc ccattttatg tgcagattat tttcagaggg tatatagaat tcaggcagct 421  
gttttcgttgt agcacattaa aaatattttc ccc 454

<210> 87  
<211> 1272  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 32..307

<221> sig\_peptide  
<222> 32..70  
<223> Von Heijne matrix  
score 4.2  
seq MLFSLSLLSNLNQ/IG

<221> polyA\_signal  
<222> 1240..1245

<221> polyA\_site  
<222> 1261..1272

<400> 87

```

gtcagggttgc accgcccttt gggtcccagag c atg ctg ttt tct ctc agc ctt      52
                               Met Leu Phe Ser Leu Ser Leu
                               -10
ctc tcc aac ctt aac caa atc ggc agc agc cac ctc gac cgc cca cac      100
Leu Ser Asn Leu Asn Gln Ile Gly Ser Ser His Leu Asp Arg Pro His
-5                               1                               5                               10
att cct ggc caa tca gct cag ctg ttt att tac caa atg tct tca caa      148
Ile Pro Gly Gln Ser Ala Gln Leu Phe Ile Tyr Gln Met Ser Ser Gln
                               15                               20                               25
caa cta cag cag cag cct tcg gct aac aaa aaa gca gga aaa atc cac      196
Gln Leu Gln Gln Gln Pro Ser Ala Asn Lys Lys Ala Gly Lys Ile His
                               30                               35                               40
aac acc ccc ttc gcc aac caa cta aat cca acg caa cat ctg gca aaa      244
Asn Thr Pro Phe Ala Asn Gln Leu Asn Pro Thr Gln His Leu Ala Lys
                               45                               50                               55
cct ttt cag caa att ctt cct ggc cgt cag tcc ggc agc ctc acc tca      292
Pro Phe Gln Gln Ile Leu Pro Gly Arg Gln Ser Gly Ser Leu Thr Ser
                               60                               65                               70
cca ttt cta gct tgc tgaaacccaa aactaatctc caagaaggag aagcttctct      347
Pro Phe Leu Ala Cys
75
cgcagccgga gcaggtcctt ttctagagat aggagaagag agagatcgct gtctcgggag      407
agaaatcaca agccgtcccg atccttctct aggtctcgta gtcgatttag gtcaaataaa      467
aggaaataga agacagtttg caagagaagt ggtgtacagg aaattacttc atttgacagg      527
agtatgtaca gaaaattcaa gttttgtttg agacttcata agcttgggtgc atttttaaga      587
tgtttttagct gttcaaatct gtttgtctct tgaaacagtg acacaaaagt gtaattctct      647
atggtttgaa atggatcata cgaggcatgt aataccaaga attgttactt tacaatgttc      707
ccttaagcaa aattgaattt gctttgaact tttagttatg cacagactga taataaacct      767
ctaaacctgc ccagcggaag tgtgtttttt tttaaattta aatacagaaa caactggcaa      827
aaattgaact aagatttact tttttttcca tagctgggat ataggctgca gctatagttg      887
aacaagcagt ctttaaaaac tgctgtgaaa cacaggccat cagggaaaac gaaatgctgc      947
actattaaat tagaggtttt tgaaaaatcc aactctcatc ctgggcagag gttgcctagt      1007
tggtatagaa tgtaagttt caagaaagtt tacctttgct ttaggtcgta agttccttat      1067
ttgattgccg tatatggata catggctgtt cgtgacattc tttatgtgca aatttgtgat      1127
ttcaaaaatg tctgccagt ttaagggtac attgtagagc cgaactttga gttactgtgc      1187
aagatttttt ttcagtctgt catttgtaat atgttttgtg agaatccttg ggattaaagt      1247
tttggttaca gattaaaaaa aaaaaa
1272

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<210> 88

<211> 804

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 114..734

<221> sig\_peptide

<222> 114..239

<223> Von Heijne matrix

score 5.2

seq LLFDLVCHEFCQS/DD

<221> polyA\_signal

<222> 768..773

<221> polyA\_site

<222> 793..804

<400> 88

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ccaacaccag gaagagtctg aagagcagcc agtgtttcgg cttgtgccct gtataacttga      60
agctgccaaa caagtacggg agttctgaaa atccagaatg gcttgatggt tac atg      116
                                     Met
cac att tta caa ctg ctt act aca gtg gat gat gga att caa gca att      164
His Ile Leu Gln Leu Leu Thr Thr Val Asp Asp Gly Ile Gln Ala Ile
-40 -35 -30
gta cat tgt cct gac act gga aaa gac att tgg aat tta ctt ttt gac      212
Val His Cys Pro Asp Thr Gly Lys Asp Ile Trp Asn Leu Leu Phe Asp
-25 -20 -15 -10
ctg gtc tgc cat gaa ttc tgc cag tct gat gat cca ccc atc att ctt      260
Leu Val Cys His Glu Phe Cys Gln Ser Asp Asp Pro Pro Ile Ile Leu
-5 1 5
caa gaa cag aaa aca gtg cta gcc tct gtt ttt tca gtg ttg tct gcc      308
Gln Glu Gln Lys Thr Val Leu Ala Ser Val Phe Ser Val Leu Ser Ala
10 15 20
atc tat gcc tca cag act gag caa gag tat cta aag ata gaa aaa gta      356
Ile Tyr Ala Ser Gln Thr Glu Gln Glu Tyr Leu Lys Ile Glu Lys Val
25 30 35
gat ctt cct cta att gac agc ctc att cgg gtc tta caa aat atg gaa      404
Asp Leu Pro Leu Ile Asp Ser Leu Ile Arg Val Leu Gln Asn Met Glu
40 45 50 55
cag tgt cag aaa aaa cca gag aac tcg gca gag tct aac aca gag gaa      452
Gln Cys Gln Lys Lys Pro Glu Asn Ser Ala Glu Ser Asn Thr Glu Glu
60 65 70
act aaa agg act gat tta acc caa gat gat ctc cac ttg aaa atc tta      500
Thr Lys Arg Thr Asp Leu Thr Gln Asp Asp Leu His Leu Lys Ile Leu
75 80 85
aag gat att tta tgt gaa ttt ctt tct aat att ttt cag gca tta aca      548
Lys Asp Ile Leu Cys Glu Phe Leu Ser Asn Ile Phe Gln Ala Leu Thr
90 95 100
aag gag acg gtg gct cag gga gta aag gaa ggc cag ttg agc aaa cag      596
Lys Glu Thr Val Ala Gln Gly Val Lys Glu Gly Gln Leu Ser Lys Gln
105 110 115
aag tgt tcc tct gca ttt caa aac ctt ctt cct ttc tat agc cct gtg      644
Lys Cys Ser Ser Ala Phe Gln Asn Leu Leu Pro Phe Tyr Ser Pro Val
120 125 130 135
gtg gaa gat ttt att aaa atc cta cgt gaa gtt gat aag gcg ctt gct      692
Val Glu Asp Phe Ile Lys Ile Leu Arg Glu Val Asp Lys Ala Leu Ala
140 145 150
gat gac ttg gaa aaa aac ttc cca agt ttg aag gtt cag act      734
Asp Asp Leu Glu Lys Asn Phe Pro Ser Leu Lys Val Gln Thr
155 160 165
taaaacctga attggaatta cttctgtaca agaaataaac tttatttttc tcaactgacaa      794
aaaaaaaaaa
804

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<210> 89

<211> 802

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 199..801

<221> polyA\_signal

<222> 780..785

<221> polyA\_site

<222> 791..802

<400> 89

agtcaccgcc	tgcttcgcac	tgagcctccc	gactcagact	ctgagtcacag	ctccgaagag	60
gaagaggaat	tcggtgtggt	tggaatcgc	tctcgctttg	ccaagggaga	ctatttacga	120
tgctgcaaga	tctgttatcc	gctctgtggt	tttgtcatcc	ttgctgcctg	tggtgtggcc	180
tggtgtggct	tggtgtgg	atg cag gtt gct ctc aag gag gat ctg gat gcc				231
		Met Gln Val Ala Leu Lys Glu Asp Leu Asp Ala				
		1 5 10				
ctc aag gaa aaa ttt cga aca atg gaa tct aat cag aaa agc tca ttc						279
Leu Lys Glu Lys Phe Arg Thr Met Glu Ser Asn Gln Lys Ser Ser Phe						
		15 20 25				
caa gaa atc ccc aaa ctt aat gaa gaa cta ctc agc aag caa aaa caa						327
Gln Glu Ile Pro Lys Leu Asn Glu Glu Leu Leu Ser Lys Gln Lys Gln						
		30 35 40				
ctt gag aag att gaa tct gga gag atg ggt ttg aac aaa gtc tgg ata						375
Leu Glu Lys Ile Glu Ser Gly Glu Met Gly Leu Asn Lys Val Trp Ile						
		45 50 55				
aac atc aca gaa atg aat aag cag att tct ctg ttg act tct gca gtg						423
Asn Ile Thr Glu Met Asn Lys Gln Ile Ser Leu Leu Thr Ser Ala Val						
		60 65 70 75				
aac cac ctc aaa gcc aat gtt aag tca gct gca gac ttg att agc ctg						471
Asn His Leu Lys Ala Asn Val Lys Ser Ala Ala Asp Leu Ile Ser Leu						
		80 85 90				
cct acc act gta gag gga ctt cag aag agt gta gct tcc att ggc aat						519
Pro Thr Thr Val Glu Gly Leu Gln Lys Ser Val Ala Ser Ile Gly Asn						
		95 100 105				
act tta aac agc gtc cat ctt gct gtg gaa gca cta cag aaa act gtg						567
Thr Leu Asn Ser Val His Leu Ala Val Glu Ala Leu Gln Lys Thr Val						
		110 115 120				
gat gaa cac aag aaa acg atg gaa tta ctg cag agt gat atg aat cag						615
Asp Glu His Lys Lys Thr Met Glu Leu Leu Gln Ser Asp Met Asn Gln						
		125 130 135				
cac ttc ttg aag gag act cct gga agc aac cag atc att ccg tca cct						663
His Phe Leu Lys Glu Thr Pro Gly Ser Asn Gln Ile Ile Pro Ser Pro						
		140 145 150 155				
tca gcc aca tca gaa ctt gac aat aaa acc cac agt gag aat ttg aaa						711
Ser Ala Thr Ser Glu Leu Asp Asn Lys Thr His Ser Glu Asn Leu Lys						
		160 165 170				
cag atg ggt gat aga tct gcc act ctg aaa aga cag tct ttg gac caa						759
Gln Met Gly Asp Arg Ser Ala Thr Leu Lys Arg Gln Ser Leu Asp Gln						
		175 180 185				
gtc acc aac aga aca gat aca gta aaa atc caa aaa aaa a						802
Val Thr Asn Arg Thr Asp Thr Val Lys Ile Gln Lys Lys Lys						
		190 195 200				

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 <212> DNA  
 <213> Homo sapiens

<220>  
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 <222> 38..1174

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 score 7.3  
 seq LLSACLVTWGLG/EP

<221> polyA\_signal  
 <222> 1452..1457

<221> polyA\_site  
 <222> 1478..1490

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 Met Pro His Ser Ser Leu  
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 cat cca tcc atc ccg tgt ccc agg ggt cac ggg gcc cag aag gca gcc 103  
 His Pro Ser Ile Pro Cys Pro Arg Gly His Gly Ala Gln Lys Ala Ala  
 -30 -25 -20  
 ttg gtt ctg ctg agt gcc tgc ctg gtg acc ctt tgg ggg cta gga gag 151  
 Leu Val Leu Leu Ser Ala Cys Leu Val Thr Leu Trp Gly Leu Gly Glu  
 -15 -10 -5 1  
 cca cca gag cac act ctc cgg tac ctg gtc ctc cac cta gcc tcc ctg 199  
 Pro Pro Glu His Thr Leu Arg Tyr Leu Val Leu His Leu Ala Ser Leu  
 5 10 15  
 cag ctg gga ctg ctg tta aac ggg gtc tgc agc ctg gct gag gag ctg 247  
 Gln Leu Gly Leu Leu Leu Asn Gly Val Cys Ser Leu Ala Glu Glu Leu  
 20 25 30  
 cgc cac atc cac tcc agg tac cgg ggc agc tac tgg agg act gtg cgg 295  
 Arg His Ile His Ser Arg Tyr Arg Gly Ser Tyr Trp Arg Thr Val Arg  
 35 40 45  
 gcc tgc ctg ggc tgc ccc ctc cgc cgt ggg gcc ctg ttg ctg ctg tcc 343  
 Ala Cys Leu Gly Cys Pro Leu Arg Arg Gly Ala Leu Leu Leu Leu Ser  
 50 55 60 65  
 atc tat ttc tac tac tcc ctc cca aat gcg gtc ggc ccg ccc ttc act 391  
 Ile Tyr Phe Tyr Tyr Ser Leu Pro Asn Ala Val Gly Pro Pro Phe Thr  
 70 75 80  
 tgg atg ctt gcc ctc ctg ggc ctc tgc cag gca ctg aac atc ctc ctg 439  
 Trp Met Leu Ala Leu Leu Gly Leu Ser Gln Ala Leu Asn Ile Leu Leu  
 85 90 95  
 ggc ctc aag ggc ctg gcc cca gct gag atc tct gca gtg tgt gaa aaa 487  
 Gly Leu Lys Gly Leu Ala Pro Ala Glu Ile Ser Ala Val Cys Glu Lys  
 100 105 110  
 ggg aat ttc aac gtg gcc cat ggg ctg gca tgg tca tat tac atc gga 535  
 Gly Asn Phe Asn Val Ala His Gly Leu Ala Trp Ser Tyr Tyr Ile Gly  
 115 120 125  
 tat ctg cgg ctg atc ctg cca gag ctc cag gcc cgg att cga act tac 583



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Tyr Leu Arg Leu Ile Leu Pro Glu Leu Gln Ala Arg Ile Arg Thr Tyr
130          135          140          145
aat cag cat tac aac aac ctg cta cgg ggt gca gtg agc cag cgg ctg      631
Asn Gln His Tyr Asn Asn Leu Leu Arg Gly Ala Val Ser Gln Arg Leu
          150          155          160
tat att ctc ctc cca ttg gac tgt ggg gtg cct gat aac ctg agt atg      679
Tyr Ile Leu Leu Pro Leu Asp Cys Gly Val Pro Asp Asn Leu Ser Met
          165          170          175
gct gac ccc aac att cgc ttc ctg gat aaa ctg ccc cag cag acc ggt      727
Ala Asp Pro Asn Ile Arg Phe Leu Asp Lys Leu Pro Gln Gln Thr Gly
          180          185          190
gac cgt gct ggc atc aag gat cgg gtt tac agc aac agc atc tat gag      775
Asp Arg Ala Gly Ile Lys Asp Arg Val Tyr Ser Asn Ser Ile Tyr Glu
          195          200          205
ctt ctg gag aac ggg cag cgg gcg ggc acc tgt gtc ctg gag tac gcc      823
Leu Leu Glu Asn Gly Gln Arg Ala Gly Thr Cys Val Leu Glu Tyr Ala
210          215          220          225
acc ccc ttg cag act ttg ttt gcc atg tca caa tac agt caa gct ggc      871
Thr Pro Leu Gln Thr Leu Phe Ala Met Ser Gln Tyr Ser Gln Ala Gly
          230          235          240
ttt agc cgg gag gat agg ctt gag cag gcc aaa ctc ttc tgc cgg aca      919
Phe Ser Arg Glu Asp Arg Leu Glu Gln Ala Lys Leu Phe Cys Arg Thr
          245          250          255
ctt gag gac atc ctg gca gat gcc cct gag tct cag aac aac tgc cgc      967
Leu Glu Asp Ile Leu Ala Asp Ala Pro Glu Ser Gln Asn Asn Cys Arg
          260          265          270
ctc att gcc tac cag gaa cct gca gat gac agc agc ttc tcg ctg tcc      1015
Leu Ile Ala Tyr Gln Glu Pro Ala Asp Asp Ser Ser Phe Ser Leu Ser
          275          280          285
cag gag gtt ctc cgg cac ctg cgg cag gag gaa aag gaa gag gtt acc      1063
Gln Glu Val Leu Arg His Leu Arg Gln Glu Glu Lys Glu Glu Val Thr
290          295          300          305
gtg ggc agc ttg aag acc tca gcg gtg ccc agt acc tcc acg atg tcc      1111
Val Gly Ser Leu Lys Thr Ser Ala Val Pro Ser Thr Ser Thr Met Ser
          310          315          320
caa gag cct gag ctc ctc ctc agt gga atg gga aag ccc ctc cct ctc      1159
Gln Glu Pro Glu Leu Leu Leu Ser Gly Met Gly Lys Pro Leu Pro Leu
          325          330          335
cgc acg gat ttc tot tgagaccag ggtcaccagg ccagagcctc cagtggctctc      1214
Arg Thr Asp Phe Ser
          340
caagcctctg gactggggggc tctcttcagt ggctgaatgt ccagcagagc tatttccttc      1274
cacagggggc cttgcaggggagggtccagg acttgacatc ttaagatgcg tcttgtcccc      1334
ttgggccagt catttccctctctgagcct cgggtgtcttc aacctgtgaa atgggatcat      1394
aatcactgcc ttacctccct cacggttggt gtgaggactg agtgtgtgga agtttttcat      1454
aaactttgga tgctagtgtta cttaaaaaaa aaaaaa      1490

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<210> 91  
 <211> 361  
 <212> DNA  
 <213> Homo sapiens

<220>  
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 <222> 26..361

<221> polyA\_site  
<222> 350..361

<400> 91

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                        Met Pro Ser Glu Gly Arg Cys Trp Glu
                        1          5
acc ttg aag gcc cta cgc agt tcc gac aaa ggt cgc ctt tgc tac tac      100
Thr Leu Lys Ala Leu Arg Ser Ser Asp Lys Gly Arg Leu Cys Tyr Tyr
10          15          20          25
cgc gac tgg ctg ctg cgg cgc gag gat gtt tta gaa gaa tgt atg tct      148
Arg Asp Trp Leu Leu Arg Arg Glu Asp Val Leu Glu Glu Cys Met Ser
          30          35          40
ctt ccc aag cta tct tct tat tct gga tgg gtg gta gag cac gtc cta      196
Leu Pro Lys Leu Ser Ser Tyr Ser Gly Trp Val Val Glu His Val Leu
          45          50          55
ccc cat atg cag gag aac caa cct ctg tct gag act tcg cca tcc tct      244
Pro His Met Gln Glu Asn Gln Pro Leu Ser Glu Thr Ser Pro Ser Ser
          60          65          70
acg tca gct tca gcc cta gat caa ccc tca ttt gtt ccc aaa tct cct      292
Thr Ser Ala Ser Ala Leu Asp Gln Pro Ser Phe Val Pro Lys Ser Pro
          75          80          85
gac gca agc tct gcc ttt tcc cca gcc tcc cct gca aca cca aat gga      340
Asp Ala Ser Ser Ala Phe Ser Pro Ala Ser Pro Ala Thr Pro Asn Gly
90          95          100          105
acc aag ggc aaa aaa aaa aaa
Thr Lys Gly Lys Lys Lys Lys
          110
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<210> 92

<211> 605

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 3..131

<221> polyA\_site

<222> 591..605

<400> 92

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ca tcc ctt ccc cag gct tta tgg ttc cag ttc ttc tac cac tct gga      47
Ser Leu Pro Gln Ala Leu Trp Phe Gln Phe Phe Tyr His Ser Gly
1          5          10          15
agc tcc cta gaa tct cct gga atg ctt aat gga cct ttc cag cac cga      95
Ser Ser Leu Glu Ser Pro Gly Met Leu Asn Gly Pro Phe Gln His Arg
          20          25          30
aat tca aga att atg act cat cgg tca gca gaa aag tgaggatacc      141
Asn Ser Arg Ile Met Thr His Arg Ser Ala Glu Lys
          35          40
ttttcctaac ctacctgctt cccctgcagt ttcttcacaa tcttactott tatatttttag      201
catatgtagc ttctcaggat gttaattctg ttctctctgt gttggtgtct gagcaccag      261
aaggtagagc caggggcact tataaaccag gagcattatt tgacaggcac ttaagaaaga      321
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cactgggtac gtaatcccag cactttggga ggctgaggcg gatggatcac atgaggtcag 381
gagttcgaga ccagcctggc cagcatgggt aaaccctgtc tctactaaaa atacaaaaat 441
tagctgggtg tggttgcaca cgcctgtaat cccagctacc tgggaggctg aggaggaga 501
atcgcttgaa cttgggaggc ggaggttgca gtgagcctag attttgccat tgcaactccag 561
cctgggtgac aagggcgaaa ctccatccca aaaaaaaaaa aaaa 605
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<210> 93
<211> 591
<212> DNA
<213> Homo sapiens
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<221> CDS
<222> 33..185
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<222> 33..80
<223> Von Heijne matrix
      score 3.7
      seq IALTLIPSMLSRA/AG
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<221> polyA_signal
<222> 570..575
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<221> polyA_site
<222> 586..591
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<400> 93
caatcttctc agcttataac cgtctttccc tt atg cta agg ata gcc ctt aca 53
                                     Met Leu Arg Ile Ala Leu Thr
                                     -15 -10
cttc atc cca tct atg ctg tca agg gct gct ggt tgg tgc tgg tac aag 101
Leu Ile Pro Ser Met Leu Ser Arg Ala Ala Gly Trp Cys Trp Tyr Lys
      -5 1 5
gag ccc act cag cag ttt tct tac ctt tgc ctg ccc tgc ctt tca tgg 149
Glu Pro Thr Gln Gln Phe Ser Tyr Leu Cys Leu Pro Cys Leu Ser Trp
      10 15 20
aat aag aaa ggc aac gtt ttg cag ctt cca aat ttc tgaagaaact 195
Asn Lys Lys Gly Asn Val Leu Gln Leu Pro Asn Phe
      25 30 35
aatctcagat tggcagttaa agtcaaaatg ttgccaaata tttattcctt ttgcctaagt 255
ttggctaccc ggttcaattg ctttttattt ttaatgtctt gactcttcag agttcgtacc 315
tcaaaagaac aatgagaaca tttgctttgc tttctgctga atccctaata tcaacaatct 375
atacctggac tgtccagttc tcctcctgtg ctatcttctc ttctatccaa gtagaatgta 435
tgccaggagc tccttccttc tagcaatttc tactaaaatg tccaagtaga atgtttcctt 495
ttacaatcaa attactgtat ttattaattt gctagaatcc agtaaatcat tttggtagct 555
ctggctgtgc tatcaataaa aagatgaaag caaaaa 591
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<210> 94
<211> 1150
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<220>  
<221> CDS  
<222> 184..915

<221> sig\_peptide  
<222> 184..237  
<223> Von Heijne matrix  
score 3.5  
seq LLGLELSEAEAG/AD

<221> polyA\_signal  
<222> 1119..1124

<221> polyA\_site  
<222> 1139..1150

<400> 94

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tttgaacagg atagtaggta tccggagtcg attgagggcc agagcaggca ctgggggttcg	120
gatacctgggc aaagtttccc acgttgaggg tctcgaggac gcctagatct ctttcccagg	180
gcc atg gcg aac ccg aag ctg ctg gga ctg gag cta agc gag gcg gag	228
Met Ala Asn Pro Lys Leu Leu Gly Leu Glu Leu Ser Glu Ala Glu	
-15 -10 -5	
gcg atc ggt gct gat tcg gcg cga ttt gag gag ctg ctg ctg cag gcc	276
Ala Ile Gly Ala Asp Ser Ala Arg Phe Glu Glu Leu Leu Leu Gln Ala	
1 5 10	
tcg aag gag ctc cag caa gcc cag aca acc aga cca gaa tcg aca caa	324
Ser Lys Glu Leu Gln Gln Ala Gln Thr Thr Arg Pro Glu Ser Thr Gln	
15 20 25	
atc cag cct cag cct ggt ttc tgc ata aag acc aac tcc tcg gaa ggg	372
Ile Gln Pro Gln Pro Gly Phe Cys Ile Lys Thr Asn Ser Ser Glu Gly	
30 35 40 45	
aag gtt ttc atc aac atc tgc cac tcc ccc tct atc cct cct ccc gcc	420
Lys Val Phe Ile Asn Ile Cys His Ser Pro Ser Ile Pro Pro Ala	
50 55 60	
gac gtg acc gag gag gag ctg ctt cag atg cta gag gag gac caa gct	468
Asp Val Thr Glu Glu Glu Leu Leu Gln Met Leu Glu Glu Asp Gln Ala	
65 70 75	
ggg ttt cgc atc ccc atg agt ctg gga gag cct cat gca gaa ctg gat	516
Gly Phe Arg Ile Pro Met Ser Leu Gly Glu Pro His Ala Glu Leu Asp	
80 85 90	
gca aaa ggc cag gga tgt acc gcc tac gac gta gct gtc aac agc gac	564
Ala Lys Gly Gln Gly Cys Thr Ala Tyr Asp Val Ala Val Asn Ser Asp	
95 100 105	
ttc tac cgg agg atg cag aac agc gat ttc ttg cgg gag ctc gtg atc	612
Phe Tyr Arg Arg Met Gln Asn Ser Asp Phe Leu Arg Glu Leu Val Ile	
110 115 120 125	
acc atc gcc agg gag ggc ctt gag gac ata tac aac ttg cag ctg aat	660
Thr Ile Ala Arg Glu Gly Leu Glu Asp Ile Tyr Asn Leu Gln Leu Asn	
130 135 140	
ccg gaa tgg cgc atg atg aag aac cgg cca ttc atg ggc tcc atc tcg	708
Pro Glu Trp Arg Met Met Lys Asn Arg Pro Phe Met Gly Ser Ile Ser	
145 150 155	
cag cag aac atc cgc tcg gag cag cgt cct cgg atc cag gag ctg ggg	756
Gln Gln Asn Ile Arg Ser Glu Gln Arg Pro Arg Ile Gln Glu Leu Gly	
160 165 170	
gac ctg tac acg ccc gcc ccc ggg aga gct gag tca ggg cct gaa aag	804

Asp	Leu	Tyr	Thr	Pro	Ala	Pro	Gly	Arg	Ala	Glu	Ser	Gly	Pro	Glu	Lys		
175						180					185						
cct	cac	ctg	aac	ctg	tgg	ctg	gaa	gcc	ccc	gac	ctc	ctc	ttg	gcc	gaa	852	
Pro	His	Leu	Asn	Leu	Trp	Leu	Glu	Ala	Pro	Asp	Leu	Leu	Leu	Ala	Glu		
190					195				200					205			
ggt	gac	ctc	ccc	aaa	ctg	gat	gga	gcc	ctg	ggg	ctg	tcg	ctg	gag	atc	900	
Val	Asp	Leu	Pro	Lys	Leu	Asp	Gly	Ala	Leu	Gly	Leu	Ser	Leu	Glu	Ile		
				210				215					220				
ggg	aga	acc	gcc	tgg	tgatgggggg	ccccagcag	ctgtatcatc	tagacgctta								955	
Gly	Arg	Thr	Ala	Trp													
				225													
tatcccgcgc	cagatcaact	ctcatgagag	caaggcagcc	ttccaccgga	agagaaagca	1015											
attaatggtg	gccatgccgc	ttctgccggt	gccttcttga	tcagggtgtc	tccttgtgct	1075											
tctgagatgt	ggagaagagg	ctgctggctt	ccctaaaagt	tgaaataaaa	gatttttgcg	1135											
tttaaaaaaa	aaaaa					1150											

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 <222> 1486..1491  
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 <222> 1504..1513

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Met	Glu	Arg	Gly	Leu	Lys	Ser	Ala	Asp	Pro	Arg	Asp	Gly	Thr	Gly	Tyr		
			-30					-25					-20				
act	ggc	tgg	gca	ggt	att	gct	gtg	ctt	tac	tta	cat	ctt	tat	gat	gta	153	
Thr	Gly	Trp	Ala	Gly	Ile	Ala	Val	Leu	Tyr	Leu	His	Leu	Tyr	Asp	Val		
			-15					-10					-5				
ttt	ggg	gac	cct	gcc	tac	cta	cag	tta	gca	cat	ggc	tat	gta	aag	caa	201	
Phe	Gly	Asp	Pro	Ala	Tyr	Leu	Gln	Leu	Ala	His	Gly	Tyr	Val	Lys	Gln		
	1			5					10								
agt	ctg	aac	tgc	tta	acc	aag	cgc	tcc	atc	acc	ttc	ctt	tgt	ggg	gat	249	
Ser	Leu	Asn	Cys	Leu	Thr	Lys	Arg	Ser	Ile	Thr	Phe	Leu	Cys	Gly	Asp		
15				20				25					30				
gca	ggc	ccc	ctg	gca	gtg	gcc	gct	gtg	cta	tat	cat	aag	atg	aac	aat	297	
Ala	Gly	Pro	Leu	Ala	Val	Ala	Ala	Val	Leu	Tyr	His	Lys	Met	Asn	Asn		
			35					40					45				
gag	aag	cag	gca	gaa	gat	tgc	atc	aca	cgg	cta	att	cac	cta	aat	aag	345	

Glu Lys Gln Ala Glu Asp Cys Ile Thr Arg Leu Ile His Leu Asn Lys	
50 55 60	
att gat cct cat gct cca aat gaa atg ctc tat ggg cga ata ggc tac	393
Ile Asp Pro His Ala Pro Asn Glu Met Leu Tyr Gly Arg Ile Gly Tyr	
65 70 75	
atc tat gct ctt ctt ttt gtc aat aag aac ttt gga gtg gaa aag act	441
Ile Tyr Ala Leu Leu Phe Val Asn Lys Asn Phe Gly Val Glu Lys Thr	
80 85 90	
cct caa agc cat att cag cag att tgt gaa aca att tta acc tct gga	489
Pro Gln Ser His Ile Gln Gln Ile Cys Glu Thr Ile Leu Thr Ser Gly	
95 100 105 110	
gaa aac cta gct agg aag aga aac ttc acg gca aag tct cca ctg atg	537
Glu Asn Leu Ala Arg Lys Arg Asn Phe Thr Ala Lys Ser Pro Leu Met	
115 120 125	
tat gaa tgg tac cag gaa tat tat gta ggg gct gct cat ggc ctg gct	585
Tyr Glu Trp Tyr Gln Glu Tyr Tyr Val Gly Ala Ala His Gly Leu Ala	
130 135 140	
gga att tat tac tac ctg atg cag ccc agc ctt caa gtg agc caa ggg	633
Gly Ile Tyr Tyr Tyr Leu Met Gln Pro Ser Leu Gln Val Ser Gln Gly	
145 150 155	
aag tta cat agt ttg gtc aag ccc agt gta gac tac gtc tgc cag ctg	681
Lys Leu His Ser Leu Val Lys Pro Ser Val Asp Tyr Val Cys Gln Leu	
160 165 170	
aaa ttc cct tct ggc aat tac cct cca tgt ata ggt gat aat cga gat	729
Lys Phe Pro Ser Gly Asn Tyr Pro Pro Cys Ile Gly Asp Asn Arg Asp	
175 180 185 190	
ctg ctt gtc cat tgg tgc cat ggc gcc cct ggg gta atc tac atg ctc	777
Leu Leu Val His Trp Cys His Gly Ala Pro Gly Val Ile Tyr Met Leu	
195 200 205	
atc cag gcc tat aag gta ttc aga gag gaa aag tat ctc tgt gat gcc	825
Ile Gln Ala Tyr Lys Val Phe Arg Glu Glu Lys Tyr Leu Cys Asp Ala	
210 215 220	
tat cag tgt gct gat gtg atc tgg caa tat ggg ttg ctg aag aag gga	873
Tyr Gln Cys Ala Asp Val Ile Trp Gln Tyr Gly Leu Leu Lys Lys Gly	
225 230 235	
tat ggg ctg tgc cac ggt tct gca ggg aat gcc tat gcc ttc ctg aca	921
Tyr Gly Leu Cys His Gly Ser Ala Gly Asn Ala Tyr Ala Phe Leu Thr	
240 245 250	
ctc tac aac ctc aca cag gac atg aag tac ctg tat agg gcc tgt aag	969
Leu Tyr Asn Leu Thr Gln Asp Met Lys Tyr Leu Tyr Arg Ala Cys Lys	
255 260 265 270	
ttt gct gaa tgg tgc tta gag tat gga gaa cat gga tgc aga aca cca	1017
Phe Ala Glu Trp Cys Leu Glu Tyr Gly Glu His Gly Cys Arg Thr Pro	
275 280 285	
gac acc cct ttc tct ctc ttt gaa gga atg gct ggg aca ata tat ttc	1065
Asp Thr Pro Phe Ser Leu Phe Glu Gly Met Ala Gly Thr Ile Tyr Phe	
290 295 300	
ctg gct gac ctg cta gtc ccc aca aaa gcc agg ttc cct gca ttt gaa	1113
Leu Ala Asp Leu Leu Val Pro Thr Lys Ala Arg Phe Pro Ala Phe Glu	
305 310 315	
ctc tgaaaggata gcatgccacc tgcaactcac tgcattgaccc tttctgtata	1166
Leu	
ttcaaaccga agctaagtgc ttccgttgct ttcaaaggaa acaaagagtc aaactgtgga	1226
cttgattttg ttagcttttt tcagaattta tctttcattc agttcccttc cattatcatt	1286
tacttttact tagaagtatc caaggaagtc ttttaacttt aattttccatt tcttctaaaa	1346
gggagagtga gtgatatgta cagtgttttg agattgtata catatatattcc agaacttgga	1406
ggaaatctta ttttaagtta tgaatataac catctgtttac tgtttctaaaa atgttttaaaa	1466

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1513

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<211> 417  
<212> DNA  
<213> Homo sapiens

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<222> 327..416  
  
<221> polyA\_site  
<222> 404..417

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tgttgttatc cttggatgta caaaaaattc agaaaatgat ctctgtagat attctgtttt 180  
attttggtca tctttagaag ttatcaggaa tgtgttttaa acaagaagag aacttttcta 240  
aggaatgata catagaaaag attttatttt aaaatgagtt gtaaagcttg tgtttctttg 300  
ttgctgcaag ctatctgccc aagtta atg caa atg gac aca ttt ttt atg tca 353  
Met Gln Met Asp Thr Phe Phe Met Ser  
1 5  
gaa aaa cac aca cac aca cac aca cat ata cac aca cac aca cga aaa 401  
Glu Lys His Thr His Thr His Thr His Ile His Thr His Thr Arg Lys  
10 15 20 25  
aca aaa aaa aaa aaa a 417  
Thr Lys Lys Lys Lys  
30

<210> 97  
<211> 603  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 63..398  
  
<221> sig\_peptide  
<222> 63..206  
<223> Von Heijne matrix  
score 4.9  
seq PSLAAGLLFGSLA/GL

<400> 97  
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aa atg cag gac act ggc tca gta gtg cct ttg cat tgg ttt ggc ttt 107  
Met Gln Asp Thr Gly Ser Val Val Pro Leu His Trp Phe Gly Phe  
-45 -40 -35  
ggc tac gca gca ctg gtt gct tct ggt ggg atc att ggc tat gta aaa 155  
Gly Tyr Ala Ala Leu Val Ala Ser Gly Gly Ile Ile Gly Tyr Val Lys  
-30 -25 -20

gca ggc agc gtg ccg tcc ctg gct gca ggg ctg ctc ttt ggc agt cta 203  
Ala Gly Ser Val Pro Ser Leu Ala Ala Gly Leu Leu Phe Gly Ser Leu  
-15 -10 -5  
gcc ggc ctg ggt gct tac cag ctg tct cag gat cca agg aac gtt tgg 251  
Ala Gly Leu Gly Ala Tyr Gln Leu Ser Gln Asp Pro Arg Asn Val Trp  
1 5 10 15  
gtt ttc cta gct aca tct ggt acc ttg gct ggc att atg gga atg agg 299  
Val Phe Leu Ala Thr Ser Gly Thr Leu Ala Gly Ile Met Gly Met Arg  
20 25 30  
ttc tac cac tct gga aaa ttc atg cct gca ggt tta att gca ggt gcc 347  
Phe Tyr His Ser Gly Lys Phe Met Pro Ala Gly Leu Ile Ala Gly Ala  
35 40 45  
agt ttg ctg atg gtc gcc aaa gtt gga gtt agt atg ttc aac aga ccc 395  
Ser Leu Leu Met Val Ala Lys Val Gly Val Ser Met Phe Asn Arg Pro  
50 55 60  
cat tagcagaagt catgttccag cttagactga tgaagaatta aaaatctgca 448  
His  
tcttccacta ttttcaatat attaagagaa ataagtgcag catttttgca tctgacattt 508  
tacctaataaa aaaagacacc aaacttggca gagaggtgga aaatcagtca tgattacaaa 568  
cctacagagg tggcgagtat gtaacacaag agctt 603

<210> 98  
<211> 522  
<212> DNA  
<213> Homo sapiens  
<220>  
<221> CDS  
<222> 2..163  
<221> polyA\_signal  
<222> 488..493  
<221> polyA\_site  
<222> 511..522

<400> 98  
c gag att gcg ggc tat ggc gcc gaa ggt ttt tcg tca gta ctg gga tat 49  
Glu Ile Ala Gly Tyr Gly Ala Glu Gly Phe Ser Ser Val Leu Gly Tyr  
1 5 10 15  
ccc cga tgg cac cga ttg cca ccg caa agc cta cag cac cac cag tat 97  
Pro Arg Trp His Arg Leu Pro Pro Gln Ser Leu Gln His His Gln Tyr  
20 25 30  
tgc cag cgt cgc tgg cct gac cgc cgc tgc cta cag agt cac act caa 145  
Cys Gln Arg Arg Trp Pro Asp Arg Arg Cys Leu Gln Ser His Thr Gln  
35 40 45  
tcc tcc ggg cac ctt cct nntgaaggag tggctaaggt tggacaatac 193  
Ser Ser Gly His Leu Pro  
50  
acgttcactg cagctgctgt cggggccgtg tttggcctca ccacctgcat cagcgcccat 253  
gtccgcgaga agcccgacga cccctgaac tacttccccg gtggctgcgc cnggaggcct 313  
gactctggga gcacgcacgc acaactacgg gattggcgcc gccgcctgcg tgtactttgg 373  
catagcggcc tccctggtca agatgggccc gctggagggc tgggaggtgt ttgcaaaacc 433  
caaggtgtga gccctgtgcc tgccgggacc tccagcctgc agaattgcgtc cagaaataaa 493  
ttctgtgtct gtgtgtgaaa aaaaaaaaaa 522



<210> 99  
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<212> DNA  
<213> Homo sapiens

<220>  
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<222> 13..465

<221> sig\_peptide  
<222> 13..75  
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score 3.9  
seq PVAVTAAVAPVLS/IN

<400> 99

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ngagtcggga aa atg gct gcg agt acn tcn atg gnc ccg gtg gct gtg acg      51
      Met Ala Ala Ser Thr Ser Met Xaa Pro Val Ala Val Thr
            -20            -15            -10
gcg gca gtg gcg cct gtc ctg tcc ata aac agc gat ttc tca gat ttg      99
Ala Ala Val Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu
            -5            1            5
cgg gaa att aaa aag caa ctg ctg ctt att gcg ggc ctt acc cgg gag      147
Arg Glu Ile Lys Lys Gln Leu Leu Leu Ile Ala Gly Leu Thr Arg Glu
      10            15            20
cgg ggc cta cta cac agt agc aaa tgg tgc gcg gag ttg gct ttc tct      195
Arg Gly Leu Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser
      25            30            35            40
ctc cct gca ttg cct cnt ggc cag ctg caa ccg cct ccg cct att aca      243
Leu Pro Ala Leu Pro Xaa Gly Gln Leu Gln Pro Pro Pro Pro Ile Thr
            45            50            55
gag gaa gat gcc cag gat atg gat gcc tat acc ctg gcc aag gcc tac      291
Glu Glu Asp Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr
            60            65            70
ttt gac gtt aaa gag tat gat cgg gca gca cat ttc ctg cat ggc tgc      339
Phe Asp Val Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys
            75            80            85
aat agc aag aaa gcc tat ttt ctg tat atg tat tcc aga tat ctg gtg      387
Asn Ser Lys Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val
            90            95            100
agg gcc att tta aaa tgt cat tct gcc ttt agt gaa aca tcc ata ttt      435
Arg Ala Ile Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe
      105            110            115            120
aga acc aat gga aaa gtt aaa tct ttt aaa tagcttagca gtgggccact      485
Arg Thr Asn Gly Lys Val Lys Ser Phe Lys
            125            130
gaatgaatgt actttataca tagcaataat aaaaaaaaga tatcataaat aaagttaaaa      545
aggatggtag agaagaaaat attccttagga atgactaaca ggataagtaa caacctgatt      605
at ttattttac tttaggttat ataaggttct tcatgcctgt gaattaatat tattgtgtaa      665
gaattaagtt aaaaagcctg ggctgacttt taaattttata aattcattta tcatgtttat      725
agtatatatta ttgtttttct ttcattggcta ttaaaaaagta tgactgtaaa ggacaatgca      785
agnaaaccaa cttaatactg tattgaataa taagtacaat ttattatttt actttgaaac      845
attatgaatt tactttccta ctttttctta gttgttatct atataaattg attaaaaaaa      905
cattttatgt acntnncatt tcctagtaca ggttgagtat cccttatttg a      956
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<210> 100  
<211> 1041  
<212> DNA  
<213> Homo sapiens

<220>  
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<222> 20..703

<221> sig\_peptide  
<222> 20..94  
<223> Von Heijne matrix  
score 3.9  
seq ATVGLLMLGVTLF/NS

<221> polyA\_signal  
<222> 1000..1005

<221> polyA\_site  
<222> 1023..1041

<400> 100

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                        Met Ser Met Ala Val Glu Thr Phe Gly Phe Phe
                        -25                -20                -15
atg gca act gtg ggg ctg ctg atg ctg ggg gtg act ctg cca aac agc      100
Met Ala Thr Val Gly Leu Leu Met Leu Gly Val Thr Leu Pro Asn Ser
                        -10                -5                1
tac tgg cga gtg tcc act gtg cac ggg aac gtc atc acc acc aac acc      148
Tyr Trp Arg Val Ser Thr Val His Gly Asn Val Ile Thr Thr Asn Thr
                        5                10                15
atc ttc gag aac ctc tgg ttt agc tgt gcc acc gac tcc ctg ggc gtc      196
Ile Phe Glu Asn Leu Trp Phe Ser Cys Ala Thr Asp Ser Leu Gly Val
                        20                25                30
tac aac tgc tgg gag ttc ccg tcc atg ctg gcc ctc tct ggg tat att      244
Tyr Asn Cys Trp Glu Phe Pro Ser Met Leu Ala Leu Ser Gly Tyr Ile
                        35                40                45                50
cag gcc tgc cgg gca ctc atg atc acc gcc atc ctc ctg ggc ttc ctc      292
Gln Ala Cys Arg Ala Leu Met Ile Thr Ala Ile Leu Leu Gly Phe Leu
                        55                60                65
ggc ctc ttg cta ggc ata gcg ggc ctg cgc tgc acc aac att ggg ggc      340
Gly Leu Leu Leu Gly Ile Ala Gly Leu Arg Cys Thr Asn Ile Gly Gly
                        70                75                80
ctg gag ctc tcc agg aaa gcc aag ctg gcg gcc acc gca ggg gcc ccc      388
Leu Glu Leu Ser Arg Lys Ala Lys Leu Ala Ala Thr Ala Gly Ala Pro
                        85                90                95
cac att ctg gcc ggt atc tgc ggg atg gtg gcc atc tcc tgg tac gcc      436
His Ile Leu Ala Gly Ile Cys Gly Met Val Ala Ile Ser Trp Tyr Ala
                        100                105                110
ttc aac atc acc cgg gac ttc ttc gac ccc ttg tac ccc gga acc aag      484
Phe Asn Ile Thr Arg Asp Phe Phe Asp Pro Leu Tyr Pro Gly Thr Lys
                        115                120                125                130
tac gag ctg ggc ccc gcc ctc tac ctg ggg tgg agc gcc tca ctg atc      532
Tyr Glu Leu Gly Pro Ala Leu Tyr Leu Gly Trp Ser Ala Ser Leu Ile
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135          140          145
tcc atc ctg ggt ggc ctc tgc ctc tgc tcc gcc tgc tgc tgc ggc tct 580
Ser Ile Leu Gly Gly Leu Cys Leu Cys Ser Ala Cys Cys Cys Gly Ser
150          155          160
gac gag gac cca gcc gcc agc gcc cgg cgg ccc tac cag gct cca gtg 628
Asp Glu Asp Pro Ala Ala Ser Ala Arg Arg Pro Tyr Gln Ala Pro Val
165          170          175
tcc gtg atg ccc gtc gcc acc tcg gac caa gaa ggc gac agc agc ttt 676
Ser Val Met Pro Val Ala Thr Ser Asp Gln Glu Gly Asp Ser Ser Phe
180          185          190
ggc aaa tac ggc aga aac gcc tac gtg tagcagctct ggcccgtggg 723
Gly Lys Tyr Gly Arg Asn Ala Tyr Val
195          200
ccccgctgtc ttcccaactgc cccaaggaga ggggacctgg ccgggggcca ttcccctata 783
gtaacctcag gggccggcca cgccccgctc ccgtagcccc gccccggcca cggccccgtg 843
tcttgactc tcatggcccc tccaggccaa gaactgctct tgggaagtcg catatctccc 903
ctctgaggct ggatccctca tcttctgacc ctgggttctg ggctgtgaag gggacggtgt 963
ccccgcacgt ttgtattgtg tataaatata ttcattaata aatgcatatt gtgaccgtta 1023
aaaaaaaaa aaaaaaaaa 1041

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<210> 101
<211> 558
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 103..294

<221> sig_peptide
<222> 103..243
<223> Von Heijne matrix
      score 5.9
      seq TWLGLLSFQNLHC/FP

<400> 101

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ttcccatggt ttagaagcat aacctgtaat gtaatgcaag tcccctaact cctggttgc 60
taacattaac ttccttaagt aataatcaat gaaagaaatt ct atg cat ggt ttt 114
                               Met His Gly Phe
                               -45
gaa ata ata tcc ttg aaa gag gaa tca cca tta gga aag gtg agt cag 162
Glu Ile Ile Ser Leu Lys Glu Glu Ser Pro Leu Gly Lys Val Ser Gln
-40          -35          -30
ggt cct ttg ttt aat gtg act agt ggc tca tca tca cca gtg acc tgg 210
Gly Pro Leu Phe Asn Val Thr Ser Gly Ser Ser Ser Pro Val Thr Trp
-25          -20          -15
ttg ggc cta ctc tcc ttc cag aac ctg cat tgc ttc cca gac ctc ccc 258
Leu Gly Leu Leu Ser Phe Gln Asn Leu His Cys Phe Pro Asp Leu Pro
-10          -5          1          5
act gag atg cct cta aga gcc aaa gga gtc aac act tgagcctagg 304
Thr Glu Met Pro Leu Arg Ala Lys Gly Val Asn Thr
10          15
gtgggctaca acaaaagatt ctaatttacc ttgcttcac taggtccagg cccaagtag 364
cttgctgaag gaacttaaaa agtagctgtt atttattgta ttgtataagc taaaaacatt 424
tatttttggt gaatcgaaac aattccatgt agcaatcttt tttctgttca cgggtgtttgt 484

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gatagaacct taaattccgc aagcatcagt tttttgaaaa aatgggaatt gaccggatag 544  
taacaggcaa agtt 558

<210> 102  
<211> 730  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 81..518

<221> sig\_peptide  
<222> 81..173  
<223> Von Heijne matrix  
score 3.9  
seq ILFHGVFYAGGFA/IV

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ctcgtcatgc tctttgtagc gtggtgcttc tgttgetcac aggacaactt gcctttgatg 60  
attttcaaga gagttgtgct atg atg tgg caa aag tat gca gga agc agg cgg 113  
Met Met Trp Gln Lys Tyr Ala Gly Ser Arg Arg  
-30 -25  
tca atg cct ctg gga gca agg atc ctt ttc cac ggt gtg ttc tat gcc 161  
Ser Met Pro Leu Gly Ala Arg Ile Leu Phe His Gly Val Phe Tyr Ala  
-20 -15 -10 -5  
ggg ggc ttt gcc att gtg tat tac ctc att caa aag ttt cat tcc agg 209  
Gly Gly Phe Ala Ile Val Tyr Tyr Leu Ile Gln Lys Phe His Ser Arg  
1 5 10  
gct tta tat tac aag ttg gca gtg gag cag ctg cag agc cat ccc gag 257  
Ala Leu Tyr Tyr Lys Leu Ala Val Glu Gln Leu Gln Ser His Pro Glu  
15 20 25  
gca cag gaa gct ctg ggc cct cct ctc aac atc cat tat ctc aag ctc 305  
Ala Gln Glu Ala Leu Gly Pro Pro Leu Asn Ile His Tyr Leu Lys Leu  
30 35 40  
atc gac agg gaa aac ttc gtg gac att gtt gat gcc aag ttg aag att 353  
Ile Asp Arg Glu Asn Phe Val Asp Ile Val Asp Ala Lys Leu Lys Ile  
45 50 55 60  
cct gtc tct gga tcc aaa tca gag ggc ctt ctc tac gtc cac tca tcc 401  
Pro Val Ser Gly Ser Lys Ser Glu Gly Leu Leu Tyr Val His Ser Ser  
65 70 75  
aga ggt ggc ccc ttt cag agg tgg cac ctt gac gag gtc ttt tta gag 449  
Arg Gly Gly Pro Phe Gln Arg Trp His Leu Asp Glu Val Phe Leu Glu  
80 85 90  
ctc aag gat ggt cag cag att cct gtg ttc aag ctc agt ggg gaa aac 497  
Leu Lys Asp Gly Gln Gln Ile Pro Val Phe Lys Leu Ser Gly Glu Asn  
95 100 105  
ggt gat gaa gtg aaa aag gag tagagacgac ccagaagacc cagcttgctt 548  
Gly Asp Glu Val Lys Lys Glu  
110 115  
ctagtccatc cttccctcat ctctaccata tggccactgg ggtggtggcc catctcagtg 608  
acagacactc ctgcaaccca gttttccagc caccagtggg atgatgggtat gtgccagcac 668  
atggtaattt tgggtgtaatt ctaacttggg cacaacgaat gctatttgtc atttttaaac 728  
tg 730

<210> 103  
<211> 1098  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 66..326

<221> polyA\_signal  
<222> 1066..1071

<221> polyA\_site  
<222> 1087..1098

<400> 103  
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ctctc atg gag ttg gct ccg aca gcc cgt ctg cca cca ggc cat ggt tcc 110  
Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly Ser  
1 5 10 15  
ttg ccc cat ggt gtc ctg gga ccc aga gca aca gga tct gtc acc cac 158  
Leu Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His  
20 25 30  
ctc tct ctt ctc ccc cag atc aag caa cgt gcc tca gag gct ttg ccc 206  
Leu Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro  
35 40 45  
gaa ttg ctt cgt cct gtc acc ccc atc acc aat ttt gag ggc agc cag 254  
Glu Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln  
50 55 60  
tct cag gac cac agt gga atc ttt ggc ctg gta aca aac ctg gaa gag 302  
Ser Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu  
65 70 75  
ctg gag gtg gac gat tgg gag ttc tgagcctctg caaactgtgc gcattctcca 356  
Leu Glu Val Asp Asp Trp Glu Phe  
80 85  
gccagggatg cagaggccac ccagaggccc ttcttgaggg cgggccacat tcccgccttc 416  
ctgggacagat tgggtagaaa ggacattctt ccaggaaagt tgactgctgg ctgattggga 476  
aagaaaatcc tggagagata cttcactgct ccaaggcttt tgagacacaa gggaatctca 536  
acaaccaggg atcaggaggg tccaaagccg acattcccag tcctgtgagc tcaggtgacc 596  
tcctccgcag aagagagatg ctgctctggc cctgggagct gaattccaag cccagggttt 656  
ggctccttaa acccgaggac cgccacctct tcccagtgtg tgcgaccagc ctcattctac 716  
ttaactttgc tctcagatgc ctcagatgct ataggtcagt gaaagggcga gtagtaagct 776  
gectgectcc ctccctcag acctctccct cataattcca gagaagggca tttctgtctt 836  
tttaagcaca gactaaggct ggaacagtcc atccttatcc ctcttctggc ttgggcccctg 896  
acacctaagt ctttcccacg gtttatgtgt gtgectcatt cctttcccac caagaatcca 956  
tcttagcgcc tcttgccagc tgccctgggt ctttctccaa gggccatcag tgtcttgctt 1016  
agcttgaggg ctttaagtct tatgctgtgt tagtttcggt gtcagaacaa attaaaattt 1076  
tcagagacgc aaaaaaaaaa aa 1098

<210> 104  
<211> 346  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 170..289

<221> sig\_peptide  
<222> 170..250  
<223> Von Heijne matrix  
score 3.6  
seq LLLLLITPSPSPL/LF

<400> 104  
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atatttcctc ctagttctta gcttgcttct gcattgattg gctttacaca actggcattt 120  
agtctgcatt acacaaatag acactaattt atttgaaca agcagcaaa atg aga act 178  
Met Arg Thr  
-25  
tta ttt ggt gca gtc agg gct cca ttt agt tcc ctc act ctg ctt cta 226  
Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr Leu Leu Leu  
-20 -15 -10  
atc acc cct tct ccc agc cct ctt cta ttt gat aga ggt ctg tcc ctc 274  
Ile Thr Pro Ser Pro Ser Pro Leu Leu Phe Asp Arg Gly Leu Ser Leu  
-5 1 5  
aga tca gca atg tct tagccctctt cctctcttcc attccttcc gttggtactc 329  
Arg Ser Ala Met Ser  
10  
atttcttcta acttttta 346

<210> 105  
<211> 685  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 36..497

<221> polyA\_signal  
<222> 650..655

<221> polyA\_site  
<222> 663..685

<400> 105  
aagttctgcg ctggctggcg gagtagcaag tggcc atg ggg agc ctc agc ggt 53  
Met Gly Ser Leu Ser Gly  
1 5  
ctg cgc ctg gca gca gga agc tgt ttt agg tta tgt gaa aga gat gtt 101  
Leu Arg Leu Ala Ala Gly Ser Cys Phe Arg Leu Cys Glu Arg Asp Val  
10 15 20  
tcc tca tct cta agg ctt acc aga agc tct gat ttg aag aga ata aat 149  
Ser Ser Ser Leu Arg Leu Thr Arg Ser Ser Asp Leu Lys Arg Ile Asn  
25 30 35  
gga ttt tgc aca aaa cca cag gaa agt ccc gga gct cca tcc cgc act 197  
Gly Phe Cys Thr Lys Pro Gln Glu Ser Pro Gly Ala Pro Ser Arg Thr

40	45	50	
tac aac aga gtg cct tta cac aaa cct acg gat tgg cag aaa aag atc			245
Tyr Asn Arg Val Pro Leu His Lys Pro Thr Asp Trp Gln Lys Lys Ile			
55	60	65	70
ctc ata tgg tca ggt cgc ttc aaa aag gaa gat gaa atc cca gag act			293
Leu Ile Trp Ser Gly Arg Phe Lys Lys Glu Asp Glu Ile Pro Glu Thr			
75	80	85	
gtc tcg ttg gag atg ctt gat gct gca aag aac aag atg cga gtg aag			341
Val Ser Leu Glu Met Leu Asp Ala Ala Lys Asn Lys Met Arg Val Lys			
90	95	100	
agc agc tat cta atg att gcc ctg acg gtg gta gga tgc atc ttc atg			389
Ser Ser Tyr Leu Met Ile Ala Leu Thr Val Val Gly Cys Ile Phe Met			
105	110	115	
gtt att gag ggc aag aag gct gcc caa aga cac gag act tta aca agc			437
Val Ile Glu Gly Lys Lys Ala Ala Gln Arg His Glu Thr Leu Thr Ser			
120	125	130	
ttg aac tta gaa aag aaa gct cgt ctg aaa gag gaa gca gct atg aag			485
Leu Asn Leu Glu Lys Lys Ala Arg Leu Lys Glu Glu Ala Ala Met Lys			
135	140	145	150
gcc aaa aca gag tagcagaggt atccgtgttg gctggatttt gaaaatccag			537
Ala Lys Thr Glu			
gaattatgtt ataacgtgcc tgtattaaaa aggatgtggt atgaggatcc atttcataaa			597
gtatgatttg cccaaacctg taccatttcc gtattttctgc cgtagaagta gaaataaatt			657
ttcttaaaaa aaaaaaaaaa aaaaaaaaaa			685

<210>	106
<211>	554
<212>	DNA
<213>	Homo sapiens

<220>	
<221>	CDS
<222>	18..320

<221>	polyA_signal
<222>	539..544

<221>	polyA_site
<222>	542..554

<400>	106	
aaccgctcgtg gggaagg atg gtg tgc gaa aaa tgt gaa aag aaa ctt ggt	50	
Met Val Cys Glu Lys Cys Glu Lys Lys Leu Gly		
1	5	10
act gtt atc act cca gat aca tgg aaa gat ggt gct agg aat acc aca	98	
Thr Val Ile Thr Pro Asp Thr Trp Lys Asp Gly Ala Arg Asn Thr Thr		
15	20	25
gaa agt ggt gga aga aag ctg aat aaa aat aaa gct ttg act tca aaa	146	
Glu Ser Gly Gly Arg Lys Leu Asn Lys Asn Lys Ala Leu Thr Ser Lys		
30	35	40
aaa gca aga ttt gat cca tat gga aag aat aag ttc tcc act tgt aga	194	
Lys Ala Arg Phe Asp Pro Tyr Gly Lys Asn Lys Phe Ser Thr Cys Arg		
45	50	55
att tgt aaa agt tct gtg cac caa cca ggt tct cat tac tgc cag ggc	242	
Ile Cys Lys Ser Ser Val His Gln Pro Gly Ser His Tyr Cys Gln Gly		

60	65	70	75	
tgt gcc tac aaa aaa ggc atc tgt gcg atg tgt ggn aaa aaa gtt ttg				290
Cys Ala Tyr Lys Lys Gly Ile Cys Ala Met Cys Gly Lys Lys Val Leu				
	80	85	90	
gat acc aaa aac tac aag caa aca tct gtc tagatgtatt gatggaattt				340
Asp Thr Lys Asn Tyr Lys Gln Thr Ser Val				
	95	100		
ctggcttttct aaatgatttt actttctgcc ttgaattttc aaggcataga tgtcaactta				400
cagaataaca tgttttaaga taattaagtt taaaccagag aatttgattg ttactcattt				460
tgctctcatg ttctaaacag caacagtgt actagtcttt tggtgtaa at gggtattttt				520
cttataagaa ttttaagaac taaaaaaaaa aaaa				554

<210> 107  
 <211> 1678  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 71..1438

<221> sig\_peptide  
 <222> 71..136  
 <223> Von Heijne matrix  
 score 3.5  
 seq AAPVAAGLGPVIS/RP

<221> polyA\_signal  
 <222> 1644..1649

<221> polyA\_site  
 <222> 1665..1678

<400> 107

ccgacttcca gaggagcget gtgcacgtgg agaagagcgg ggactcggcg accctgcctt	60
cccgaccctc atg ttc gaa gag cct gag tgg gcc gag gcg gcc cca gta	109
Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val	
-20 -15 -10	
gcc gcg ggc ctt ggg ccc gta atc tca cga cct ccg cct gcg gcc tcc	157
Ala Ala Gly Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser	
-5 1 5	
tcg caa aac aag ggc tcc aag cgc cgc cag ctc ttg gcc aca tta cgg	205
Ser Gln Asn Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg	
10 15 20	
gcc cta gag gca gca tct ctt tcc cag cat ccc ccc agc cta tgt ata	253
Ala Leu Glu Ala Ala Ser Leu Ser Gln His Pro Pro Ser Leu Cys Ile	
25 30 35	
agt gac tct gag gag gag gag gag gaa agg aag aag aaa tgc ccc aaa	301
Ser Asp Ser Glu Glu Glu Glu Glu Arg Lys Lys Lys Cys Pro Lys	
40 45 50 55	
aag gca tca ttt gcc agt gcc tct gct gaa gta ggg aag aaa ggg aag	349
Lys Ala Ser Phe Ala Ser Ala Ser Ala Glu Val Gly Lys Lys Gly Lys	
60 65 70	
aag aaa tgt caa aaa cag ggc cca cct tgc agt gac tct gag gaa gaa	397
Lys Lys Cys Gln Lys Gln Gly Pro Pro Cys Ser Asp Ser Glu Glu Glu	



gta gaa agg aag aag aaa tgc cac aaa cag gct ctt gtt ggc agt gac	445
Val Glu Arg Lys Lys Lys Cys His Lys Gln Ala Leu Val Gly Ser Asp	
90 95 100	
tct gct gaa gat gag aaa aga aag agg aaa tgc cag aaa cat gcc cct	493
Ser Ala Glu Asp Glu Lys Arg Lys Arg Lys Cys Gln Lys His Ala Pro	
105 110 115	
ata aat tca gcc cag cac ctg gac aat gtt gac caa aca ggt ccc aaa	541
Ile Asn Ser Ala Gln His Leu Asp Asn Val Asp Gln Thr Gly Pro Lys	
120 125 130 135	
gcc tgg aag ggt agt act aca aat gat cca cca aag caa agc cct ggg	589
Ala Trp Lys Gly Ser Thr Thr Asn Asp Pro Pro Lys Gln Ser Pro Gly	
140 145 150	
tcc act tcc cct aaa ccc cct cat aca tta agc cgc aag cag tgg cgg	637
Ser Thr Ser Pro Lys Pro Pro His Thr Leu Ser Arg Lys Gln Trp Arg	
155 160 165	
aac cgg caa aag aat aag aga aga tgt aag aac aag ttt cag cca cct	685
Asn Arg Gln Lys Asn Lys Arg Arg Cys Lys Asn Lys Phe Gln Pro Pro	
170 175 180	
cag gtg cca gac cag gcc cca gct gag gcc ccc aca gag aag aca gag	733
Gln Val Pro Asp Gln Ala Pro Ala Glu Ala Pro Thr Glu Lys Thr Glu	
185 190 195	
gtg tct cct gtt ccc agg aca gac agc cat ggg gct cgg gca ggg gct	781
Val Ser Pro Val Pro Arg Thr Asp Ser His Gly Ala Arg Ala Gly Ala	
200 205 210 215	
ttg cga gcc cgc atg gca cag cgg ctg gat ggg gcc cga ttt cgc tac	829
Leu Arg Ala Arg Met Ala Gln Arg Leu Asp Gly Ala Arg Phe Arg Tyr	
220 225 230	
ctc aat gaa cag ttg tac tca ggg ccc agc agt gct gca cag cgt ctc	877
Leu Asn Glu Gln Leu Tyr Ser Gly Pro Ser Ser Ala Ala Gln Arg Leu	
235 240 245	
ttc cag gaa gac cct gag gct ttt ctt ctc tac cac cgc ggc ttc cag	925
Phe Gln Glu Asp Pro Glu Ala Phe Leu Leu Tyr His Arg Gly Phe Gln	
250 255 260	
agc caa gtg aag aag tgg cca ctg cag cca gtg gac cgc atc gcc agg	973
Ser Gln Val Lys Lys Trp Pro Leu Gln Pro Val Asp Arg Ile Ala Arg	
265 270 275	
gat ctt cgc cag cgg cct gca tcc cta gtg gtg gct gac ttc ggc tgt	1021
Asp Leu Arg Gln Arg Pro Ala Ser Leu Val Val Ala Asp Phe Gly Cys	
280 285 290 295	
ggg gat tgc cgc ttg gct tca agt atc cgg aac cct gtg cat tgc ttt	1069
Gly Asp Cys Arg Leu Ala Ser Ser Ile Arg Asn Pro Val His Cys Phe	
300 305 310	
gac ttg gct tct ctg gac cct agg gtc act gtg tgt gac atg gcc cag	1117
Asp Leu Ala Ser Leu Asp Pro Arg Val Thr Val Cys Asp Met Ala Gln	
315 320 325	
gtt cct ttg gag gat gag tct gtg gat gtg gct gtg ttt tgc ctt tca	1165
Val Pro Leu Glu Asp Glu Ser Val Asp Val Ala Val Phe Cys Leu Ser	
330 335 340	
ctg atg gga acc aac atc agg gac ttc cta gag gag gca aat aga gta	1213
Leu Met Gly Thr Asn Ile Arg Asp Phe Leu Glu Ala Asn Arg Val	
345 350 355	
ctg aag cca ggg ggt ctc ctg aaa gtg gct gag gtc agc agc cgc ttt	1261
Leu Lys Pro Gly Gly Leu Leu Lys Val Ala Glu Val Ser Ser Arg Phe	
360 365 370 375	
gag gat gtt cga acc ttt ctg cgg gct gtg acc aag cta ggc ttc aag	1309
Glu Asp Val Arg Thr Phe Leu Arg Ala Val Thr Lys Leu Gly Phe Lys	

```

      380              385              390
att gtc tcc aag gac ctg acc aac agc cat ttc ttc ttg ttt gat ttc   1357
Ile Val Ser Lys Asp Leu Thr Asn Ser His Phe Phe Leu Phe Asp Phe
      395              400              405
caa aag act ggg ccc cct ctg gta ggg ccc aag gct cag ctt tca ggc   1405
Gln Lys Thr Gly Pro Pro Leu Val Gly Pro Lys Ala Gln Leu Ser Gly
      410              415              420
ctg cag ctt cag cca tgt ctc tac aag cgc agg tgacctctgg atcttccttg   1458
Leu Gln Leu Gln Pro Cys Leu Tyr Lys Arg Arg
      425              430
agagggggagg cagatctcaa actccaggct cagaactgtg aagactgttt ccggcctggc   1518
tgtgagccaa gacctgggtc ctggtggacc ctgaggacaa agtgtgataa aacctctggc   1578
tcagacttgc tctactgaag gcttcttggt tataagatgc ataaagtcac tggggctagc   1638
taaacaataa agagttttatt gtgaggaaaa aaaaaaaaaa   1678
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<210> 108  
<211> 494  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 25..318  
  
<221> sig\_peptide  
<222> 25..75  
<223> Von Heijne matrix  
score 7.4  
seq FFLLLQFFFLRIDG/VL  
  
<221> polyA\_signal  
<222> 452..457  
  
<221> polyA\_site  
<222> 482..494

```

<400> 108
aggctgagtg tgaagattag agta atg cct tct agc ttt ttc ctg ctg ttg   51
                        Met Pro Ser Ser Phe Phe Leu Leu Leu
                        -15              -10
cag ttt ttc ttg aga att gat ggg gtg ctt atc aga atg aat gac acg   99
Gln Phe Phe Leu Arg Ile Asp Gly Val Leu Ile Arg Met Asn Asp Thr
      -5              1              5
aga ctt tac cat gag gct gac aag acc tac atg tta cga gaa tat acg   147
Arg Leu Tyr His Glu Ala Asp Lys Thr Tyr Met Leu Arg Glu Tyr Thr
      10              15              20
tca cga gaa agc aaa att tct agt ttg atg cat gtt cca cct tcc ctc   195
Ser Arg Glu Ser Lys Ile Ser Ser Leu Met His Val Pro Pro Ser Leu
      25              30              35              40
ttc acg gaa cct aat gaa ata tcc cag tat tta cca ata aag gaa gca   243
Phe Thr Glu Pro Asn Glu Ile Ser Gln Tyr Leu Pro Ile Lys Glu Ala
      45              50              55
gtt tgt gag aag cta ata ttt cca gaa aga att gat cct aac cca gca   291
Val Cys Glu Lys Leu Ile Phe Pro Glu Arg Ile Asp Pro Asn Pro Ala
      60              65              70
```

gac tca caa aaa agt aca caa gtg gaa taaaatgtga tacaacatat 338  
Asp Ser Gln Lys Ser Thr Gln Val Glu  
75 80

```
actcactatg gaatctgact ggacaccttg gctatttgta aggggttatt tttattatga 398
gaattaattg ccttgtttat gtacagattt tctgtagcct taaaggaaaa aaaaataaag 458
atcgttacag gcaggtttca ctcaaaaaaaaa aaaaac 494
```

```
<210> 109
<211> 714
<212> DNA
<213> Homo sapiens
```

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<220>
<221> CDS
<222> 84..332
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```
<221> sig_peptide
<222> 84..170
<223> Von Heijne matrix
      score 5.2
      seq PCYYLGLFORALA/SV
```

```
<221> polyA_site  
<222> 702..714
```

```

X400> 109
cttatctctt ctgctggctg ggctcaatgc cgcgggtgag cgttcggccg aggctgctcc      60
tacccttgag tgatgtgcct tga atg acg ctg ctt tca ttc gct gct ttc acg      113
                Met Thr Leu Leu Ser Phe Ala Ala Phe Thr
                        -25                        -20

```

gct gct ttc tcc gtc ctc ccc tgt tac tac ctt ggg ctg ttt cag cgg 161  
Ala Ala Phe Ser Val Leu Pro Cys Tyr Tyr Leu Gly Leu Phe Gln Arg  
-15 -10 -5

gcg ctc gcg tgc gtc ttc gac cca ctt tgc gtt tgt tca cgt gtg ctc 209  
Ala Leu Ala Ser Val Phe Asp Pro Leu Cys Val Cys Ser Arg Val Leu  
1 5 10

ccg aca cct gta tgt acc ttg gtc gca aca caa gcc gaa aaa ata tta 257  
Pro Thr Pro Val Cys Thr Leu Val Ala Thr Gln Ala Glu Lys Ile Leu  
15 20 25

gag aat ggg ccc tgt cca acc aag gag gcg gcc cag ctt gtc ggg aag 305  
Glu Asn Gly Pro Cys Pro Thr Lys Glu Ala Ala Gln Leu Val Gly Lys  
30 35 40 45

ggc agc gtt tcc gcc aga aat gct tcg tgaaaggcac ttgagggacc 352  
Gly Ser Val Ser Ala Arg Asn Ala Ser  
50

ttagcagcat	cctcaacagg	ccttgtaggg	aatgccagaa	gaagcagtc	ttggccggggc	412
ggggtggctc	atgcctgtgg	tcccagcact	ttgggaggcc	ggggcgggcg	gatcacctga	472
ggtcgggagg	tccagaccag	cctgaccgac	atggagaaac	cccgtctnta	ctagaaatac	532
aaaactagcc	gggtgtggtg	gcgcatgcct	gtagtccag	ctactcggga	gggtgaggca	592
ggagacgttc	ttgaacccgg	gaggcggagt	ttgtggtgag	ccgagatcgc	gccattgcac	652
tccagcctgg	gcatgccaa	agcgaaactc	cgtctcaaaa	aaaaaaaaaga	aaaaaaaaaaa	712
aa						714

<210> 110  
 <211> 805  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 32..718

<221> sig\_peptide  
 <222> 32..100  
 <223> Von Heijne matrix  
 score 7.4  
 seq VLLLAALPPVLLP/GA

<221> polyA\_signal  
 <222> 770..775

<221> polyA\_site  
 <222> 793..805

<400> 110

cctcttttcag cccgggatcg ccccagcagg g atg ggc gac aag atc tgg ctg	52
Met Gly Asp Lys Ile Trp Leu	
-20	
ccc ttc ccc gtg ctc ctt ctg gcc gct ctg cct ccg gtg ctg ctg cct	100
Pro Phe Pro Val Leu Leu Leu Ala Ala Leu Pro Pro Val Leu Leu Pro	
-15 -10 -5	
ggg gcg gcc ggc ttc aca cct tcc ctc gat agc gac ttc acc ttt acc	148
Gly Ala Ala Gly Phe Thr Pro Ser Leu Asp Ser Asp Phe Thr Phe Thr	
5 10 15	
ctt ccc gcc ggc cag aag gag tgc ttc tac cag ccc atg ccc ctg aag	196
Leu Pro Ala Gly Gln Lys Glu Cys Phe Tyr Gln Pro Met Pro Leu Lys	
20 25 30	
gcc tcg ctg gag atc gag tac caa gtt tta gat gga gca gga tta gat	244
Ala Ser Leu Glu Ile Glu Tyr Gln Val Leu Asp Gly Ala Gly Leu Asp	
35 40 45	
att gat ttc cat ctt gcc tct cca gaa ggc aaa acc tta gtt ttt gaa	292
Ile Asp Phe His Leu Ala Ser Pro Glu Gly Lys Thr Leu Val Phe Glu	
50 55 60	
caa aga aaa tca gat gga gtt cac act gta gag act gaa gtt ggt gat	340
Gln Arg Lys Ser Asp Gly Val His Thr Val Glu Thr Glu Val Gly Asp	
65 70 75 80	
tac atg ttc tgc ttt gac aat aca ttc agc acc att tct gag aag gtg	388
Tyr Met Phe Cys Phe Asp Asn Thr Phe Ser Thr Ile Ser Glu Lys Val	
85 90 95	
att ttc ttt gaa tta atc ctg gat aat atg gga gaa cag gca caa gaa	436
Ile Phe Phe Glu Leu Ile Leu Asp Asn Met Gly Glu Gln Ala Gln Glu	
100 105 110	
caa gaa gat tgg aag aaa tat att act ggc aca gat ata ttg gat atg	484
Gln Glu Asp Trp Lys Lys Tyr Ile Thr Gly Thr Asp Ile Leu Asp Met	
115 120 125	
aaa ctg gaa gac atc ctg gaa tcc atc agc agc atc aag tcc aga cta	532
Lys Leu Glu Asp Ile Leu Glu Ser Ile Ser Ser Ile Lys Ser Arg Leu	
130 135 140	
agc aaa agt ggg cac ata caa att ctg ctt aga gca ttt gaa gct cgt	580
Ser Lys Ser Gly His Ile Gln Ile Leu Leu Arg Ala Phe Glu Ala Arg	

```
145          150          155          160
gat cga aac ata caa gaa agc aac ttt gat aga gtc aat ttc tgg tct      628
Asp Arg Asn Ile Gln Glu Ser Asn Phe Asp Arg Val Asn Phe Trp Ser
          165          170          175
atg gtt aat tta gtg gtc atg gtg gtg gtg tca gcc att caa gtt tat      676
Met Val Asn Leu Val Val Met Val Val Val Ser Ala Ile Gln Val Tyr
          180          185          190
atg ctg aag agt ctg ttt gaa gat aag agg aaa agt aga act      718
Met Leu Lys Ser Leu Phe Glu Asp Lys Arg Lys Ser Arg Thr
          195          200          205
taaaactcca aactagagta cgtaacattg aaaaatgagg cataaaaatg caataaactg      778
ttacagtcaa gaccaaaaaa aaaaaaa      805
```

```
<210> 111
<211> 787
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 26..481
<221> sig_peptide
<222> 26..88
<223> Von Heijne matrix
      score 4.4
      seq AVASSFFFCASLFS/AV
<221> polyA_signal
<222> 755..760
<221> polyA_site
<222> 775..787
```

```
<400> 111
gacagcctgg ataaaggctc acttg atg gct cag ttg gga gca gtt gtg gct      52
                        Met Ala Gln Leu Gly Ala Val Val Ala
                        -20                        -15
gtg gct tcc agt ttc ttt tgt gca tct ctc ttc tca gct gtg cac aag      100
Val Ala Ser Ser Phe Phe Cys Ala Ser Leu Phe Ser Ala Val His Lys
      -10      -5      1
ata gaa gag gga cat att ggg gta tat tac aga ggc ggt gcc ctg ctg      148
Ile Glu Glu Gly His Ile Gly Val Tyr Tyr Arg Gly Gly Ala Leu Leu
5      10      15      20
act tcg acc agc ggc cct ggt ttc cat ctc atg ctc cct ttc atc aca      196
Thr Ser Thr Ser Gly Pro Gly Phe His Leu Met Leu Pro Phe Ile Thr
      25      30      35
tca tat aag tct gtg cag acc aca ctc cag aca gat gag gtg aag aat      244
Ser Tyr Lys Ser Val Gln Thr Thr Leu Gln Thr Asp Glu Val Lys Asn
      40      45      50
gta cct tgt ggg act agt ggt ggt gtg atg atc tac ttt gac aga att      292
Val Pro Cys Gly Thr Ser Gly Gly Val Met Ile Tyr Phe Asp Arg Ile
      55      60      65
gaa gtg gtg aac ttc ctg gtc ccg aac gca gtg cat gat ata gtg aag      340
Glu Val Val Asn Phe Leu Val Pro Asn Ala Val His Asp Ile Val Lys
```

```

70          75          80
aac tat act gct gac tat gac aag gcc ctc atc ttc aac aag atc cac      388
Asn Tyr Thr Ala Asp Tyr Asp Lys Ala Leu Ile Phe Asn Lys Ile His
85          90          95          100
cac gaa ctg aac cag ttc tgc agt gtg cac acg ctt caa gag gtc tac      436
His Glu Leu Asn Gln Phe Cys Ser Val His Thr Leu Gln Glu Val Tyr
          105          110          115
att gag ctg ttt gga ctg gaa aat gat ttt tcc cag gaa tct tca      481
Ile Glu Leu Phe Gly Leu Glu Asn Asp Phe Ser Gln Glu Ser Ser
          120          125          130
taaaagggac cctgagcaag aacatttttc atagcagaca ggaggactca tccacatcgc      541
cagcaatcat aattaagcaa accgcctttt gcaccatttta agatttagga aatcatccaa      601
attactttta atgtttctgc agtagaaaat gaatctaaat tcattttata gggttttag      661
tcttttatct gttttggatt cactgtgctt ttaagaaaaa gtttggtaaat ttgccgttga      721
tttttctttt taacctcaaa ctaatagaat tttataaaat attaattttc tccaaaaaaa      781
aaaaaa

```

```

<210> 112
<211> 569
<212> DNA
<213> Homo sapiens

```

```

<220>
<221> CDS
<222> 26..562
<221> sig_peptide
<222> 26..187
<223> Von Heijne matrix
      score 4.1
      seq AVVAAAARTGSEA/RV

```

```

K400> 112
agaaacaggt ctgggctaca aaagt atg gcc gct tct gag gcg gcg gtg gtg      52
                               Met Ala Ala Ser Glu Ala Ala Val Val
                               -50
tct tcg ccg tct ttg aaa aca gac aca tcc cct gtc ctt gaa act gca      100
Ser Ser Pro Ser Leu Lys Thr Asp Thr Ser Pro Val Leu Glu Thr Ala
-45          -40          -35          -30
gga acg gtc gca gca atg gct gcg acc ccg tca gca agg gct gca gcc      148
Gly Thr Val Ala Ala Met Ala Ala Thr Pro Ser Ala Arg Ala Ala Ala
          -25          -20          -15
gcg gtg gtt gcg gcc gcg gcc agg acc gga tcc gaa gcc agg gtc tcc      196
Ala Val Val Ala Ala Ala Ala Arg Thr Gly Ser Glu Ala Arg Val Ser
          -10          -5          1
aag gcc gct ttg gct acc aag ctg ctg tcc ttg agc ggc gtg ttc gcc      244
Lys Ala Ala Leu Ala Thr Lys Leu Leu Ser Leu Ser Gly Val Phe Ala
5          10          15
gtg cac aag ccc aaa ggg ccc act tca gcc gag ctg ctg aat cgg ttg      292
Val His Lys Pro Lys Gly Pro Thr Ser Ala Glu Leu Leu Asn Arg Leu
20          25          30          35
aag gag aag ctg ctg gca gaa gct gga atg cct tct cca gaa tgg acc      340
Lys Glu Lys Leu Leu Ala Glu Ala Gly Met Pro Ser Pro Glu Trp Thr
          40          45          50
aag agg aaa aag cag act ttg aaa att ggg cat gga ggg act cta gac      388

```

Lys	Arg	Lys	Lys 55	Gln	Thr	Leu	Lys	Ile 60	Gly	His	Gly	Gly	Thr 65	Leu	Asp	
agc	gca	gcc	cga	gga	gtt	ctg	gtt	gtt	gga	att	gga	agc	gga	aca	aaa	436
Ser	Ala	Ala	Arg	Gly	Val	Leu	Val	Val	Gly	Ile	Gly	Ser	Gly	Thr	Lys	
		70					75					80				
atg	ttg	acc	agt	atg	ttg	tca	ggg	tcc	aag	agg	tat	act	gcc	att	gga	484
Met	Leu	Thr	Ser	Met	Leu	Ser	Gly	Ser	Lys	Arg	Tyr	Thr	Ala	Ile	Gly	
						90					95					
gaa	ctg	ggg	aaa	gct	act	gat	aca	cta	gat	tct	acg	ggg	aag	gta	aca	532
Glu	Leu	Gly	Lys	Ala	Thr	Asp	Thr	Leu	Asp	Ser	Thr	Gly	Lys	Val	Thr	
100					105					110						
gaa	gaa	aaa	cct	tac	ggt	atg	aac	ctc	atc	taagtag					115	
Glu	Glu	Lys	Pro	Tyr	Gly	Met	Asn	Leu	Ile							569
				120					125							

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<210> 113
<211> 893
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 4..810
<221> sig_peptide
<222> 4..279
<223> Von Heijne matrix
      score 6.8
      seq AVMLYLTWRSCSRA/IP
<221> polyA_signal
<222> 858..863
<221> polyA_site
<222> 881..893
<400> 113

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gcc	atg	atc	acg	cac	gtc	acc	ctg	gaa	gat	gcc	ctg	tcc	aac	gtg	gac	48
	Met	Ile	Thr	His	Val	Thr	Leu	Glu	Asp	Ala	Leu	Ser	Asn	Val	Asp	
			-90					-85					-80			
ctg	ctt	gaa	gag	ctt	ccc	ctc	ccc	gac	cag	cag	cca	tgc	atc	gag	cct	96
Leu	Leu	Glu	Glu	Leu	Pro	Leu	Pro	Asp	Gln	Gln	Pro	Cys	Ile	Glu	Pro	
			-75				-70					-65				
cca	cct	tcc	tcc	atc	atg	tac	cag	gct	aac	ttt	gac	aca	aac	ttt	gag	144
Pro	Pro	Ser	Ser	Ile	Met	Tyr	Gln	Ala	Asn	Phe	Asp	Thr	Asn	Phe	Glu	
		-60				-55					-50					
gac	agg	aat	gca	ttt	gtc	acg	ggc	att	gca	agg	tac	att	gag	cag	gct	192
Asp	Arg	Asn	Ala	Phe	Val	Thr	Gly	Ile	Ala	Arg	Tyr	Ile	Glu	Gln	Ala	
				-40						-35					-30	
aca	gtc	cac	tcc	agc	atg	aat	gag	atg	ctg	gag	gaa	gga	cat	gag	tat	240
Thr	Val	His	Ser	Ser	Met	Asn	Glu	Met	Leu	Glu	Glu	Gly	His	Glu	Tyr	
				-25					-20				-15			
gcg	gtc	atg	ctg	tac	acc	tgg	cgc	agc	tgt	tcc	cgg	gcc	att	ccc	cag	288
Ala	Val	Met	Leu	Tyr	Thr	Trp	Arg	Ser	Cys	Ser	Arg	Ala	Ile	Pro	Gln	
			-10					-5					1			

```

gtg aaa tgc aac gag cag ccc aac cga gta gag atc tat gag aag aca      336
Val Lys Cys Asn Glu Gln Pro Asn Arg Val Glu Ile Tyr Glu Lys Thr
  5              10              15
gta gag gtg ctg gag ccg gag gtc acc aag ctc atg aag ttc atg tat      384
Val Glu Val Leu Glu Pro Glu Val Thr Lys Leu Met Lys Phe Met Tyr
 20              25              30              35
ttt cag cgc aag gcc atc gag cgg ttc tgc agc gag gtg aag cgg ctg      432
Phe Gln Arg Lys Ala Ile Glu Arg Phe Cys Ser Glu Val Lys Arg Leu
              40              45              50
tgc cat gcc gag cgc agg aag gac ttt gtc tct gag gcc tac ctc ctg      480
Cys His Ala Glu Arg Arg Lys Asp Phe Val Ser Glu Ala Tyr Leu Leu
              55              60              65
acc ctt ggc aag ttc atc aac atg ttt gct gtc ctg gat gag cta aag      528
Thr Leu Gly Lys Phe Ile Asn Met Phe Ala Val Leu Asp Glu Leu Lys
              70              75              80
aac atg aag tgc agc gtc aag aat gac cac tcc gcc tac aag agg gca      576
Asn Met Lys Cys Ser Val Lys Asn Asp His Ser Ala Tyr Lys Arg Ala
              85              90              95
gca cag ttc ctg cgg aag atg gca gat ccc cag tct atc cag gag tcg      624
Ala Gln Phe Leu Arg Lys Met Ala Asp Pro Gln Ser Ile Gln Glu Ser
100              105              110              115
cag aac ctt tcc atg ttc ctg gcc aac cac aac agg atc acc cag tgt      672
Gln Asn Leu Ser Met Phe Leu Ala Asn His Asn Arg Ile Thr Gln Cys
              120              125              130
ctc cac cag caa ctt gaa gtg atc cca ggc tat gag gag ctg ctg gct      720
Leu His Gln Gln Leu Glu Val Ile Pro Gly Tyr Glu Glu Leu Leu Ala
              135              140              145
gac att gtc aac atc tgt gtg gat tac tac gag aac aag atg tac ctg      768
Asp Ile Val Asn Ile Cys Val Asp Tyr Tyr Glu Asn Lys Met Tyr Leu
              150              155              160
act ccc agt gag aaa cat atg ctc ctc aag gta aaa ctc ccc      810
Thr Pro Ser Glu Lys His Met Leu Leu Lys Val Lys Leu Pro
              165              170              175
ctgaggccgca cccatggagc ctgggcttac cctctcacct tcttcttatt aaaaatccgt      870
ttttaaaaaaac aaaaaaaaaa aaa

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<210> 114

<211> 1475

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 55..459

<221> sig\_peptide

<222> 55..120

<223> Von Heijne matrix

score 7.2

seq GLWLALVDGLVRS/SP

<221> polyA\_signal

<222> 1444..1449

<221> polyA\_site



<222> 1462..1475

<400> 114

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cagttccgca gctacgtgtg ggaccgctg ctgatcctgt cgcagatcgt cctc atg      57
                                     Met
cag acc gtg tat tac ggc tcg ctg ggc ctg tgg ctg gcg ctg gtg gac      105
Gln Thr Val Tyr Tyr Gly Ser Leu Gly Leu Trp Leu Ala Leu Val Asp
   -20                               -15               -10
ggg cta gtg cga agc agc ccc tcg ctg gac cag atg ttc gac gcc gag      153
Gly Leu Val Arg Ser Ser Pro Ser Leu Asp Gln Met Phe Asp Ala Glu
   -5                               1               5               10
atc ctg ggc ttt tcc acc cct cca ggc cgg ctc tcc atg atg tcc ttc      201
Ile Leu Gly Phe Ser Thr Pro Pro Gly Arg Leu Ser Met Met Ser Phe
          15                               20               25
atc ttc aac gcc ctc acc tgt gcc ctg ggc ttg ctg tac ttc atc cgg      249
Ile Phe Asn Ala Leu Thr Cys Ala Leu Gly Leu Leu Tyr Phe Ile Arg
          30                               35               40
cga gga aag cag tgt ctg gat ttc act gtc act gtc cat ttc ttt cac      297
Arg Gly Lys Gln Cys Leu Asp Phe Thr Val Thr Val His Phe Phe His
          45                               50               55
ctc ctg ggc tgc tgg ttc tac agc tcc cgt ttc ccc tcg gcg ctg acc      345
Leu Leu Gly Cys Trp Phe Tyr Ser Ser Arg Phe Pro Ser Ala Leu Thr
          60                               65               70               75
tgg tgg ctg gtc caa gcc gtg tgc att gca ctc atg gct gtc atc ggg      393
Trp Trp Leu Val Gln Ala Val Cys Ile Ala Leu Met Ala Val Ile Gly
          80                               85               90
gag tac ctg tgc atg cgg acg gag ctc aag gag ata ccc ctc aac tca      441
Glu Tyr Leu Cys Met Arg Thr Glu Leu Lys Glu Ile Pro Leu Asn Ser
          95                               100              105
gcc cct aaa tcc aat gtc tagaatcagg ccctttggac atcccgtga      489
Ala Pro Lys Ser Asn Val
          110
cacttggggc ccttaacacc ttgggctgct cagaccctcc agatgaggtc cagcccagat      549
ctgagaggaa ccttggaat gtgaagtctc tgttgggtgtg ggagagatag tgagggcctg      609
tcaaagaagg caggtagcag tcagcatgac agctgcaaga atgacctctg tctgttgaag      669
cccttggtatc tgagagggtca ggaaggggac ctctttgagg gtaataacat aattggaacc      729
atgccactct tgagccacaa tacctgtcac cagcctgttg ttttaagaga gaaaaaaaaat      789
caaggatatac tgattggagc aaaccacttc tttagtcac tgtcttacct ccctgggaca      849
gctgttacct ttgcagtgtt gccgaatcac agcagttacc tttgcaatgt tgccgaatca      909
cagcagttct gttggagaaa cgcttggttt ccggatccag agccacagaa agaaatgtag      969
gtgtgaagta ttaggctgct gtcagggaga ggatggcaga tggaggcatc aagcacaagg      1029
aaaatgcaca acctgtgccc tgttatacac acgttcatgt gcgccaaga acctatgact      1089
ttcttccagt tcttctacc aggtcccat cctgctgcca gctctcaaca tagcaggcca      1149
taggaccagc agaagaatcc cagtgttgct caaagtctga ccatcataaa gacactgcct      1209
gtcttctagg aatgaccagg caccagctc ccactggact ccaatttttt ttctgcctt      1269
atttagaatt ctttggcggg aagggtatga tgggttccca gagacaagaa gcccaacctt      1329
ctggcctggg ctgtgctgat agtgctgagg gagataggaa tttgctgcta agatttttct      1389
ttggggtgga gtttctctct tgaggggctt gcagctatcc ttctgtgta taaaaataca      1449
gtattttcca tgaaaaaaaa aaaaaa
                                     1475

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<210> 115

<211> 321

<212> DNA

<213> Homo sapiens

<220>  
<221> CDS  
<222> 48..248

<221> sig\_peptide  
<222> 48..161  
<223> Von Heijne matrix  
score 6.3  
seq LVFALVTAVCCCLA/DG

<221> polyA\_signal  
<222> 283..288

<221> polyA\_site  
<222> 308..321

<400> 115  
gctgagaaga gttgagggaa agtgctgctg ctgggtctgc agacgcg atg aat aac 56  
Met Asn Asn  
gtg cag ccg aaa ata aaa cat cgc ccc ttc tgc ttc agt gtg aaa ggc 104  
Val Gln Pro Lys Ile Lys His Arg Pro Phe Cys Phe Ser Val Lys Gly  
-35 -30 -25 -20  
cac gtg aag atg ctg cgg ctg gtg ttt gca ctt gtg aca gca gta tgc 152  
His Val Lys Met Leu Arg Leu Val Phe Ala Leu Val Thr Ala Val Cys  
-15 -10 -5  
tgt ctt gcc gac ggg gcc ctt att tac cgg aag ctt ctg ttc aat ccc 200  
Cys Leu Ala Asp Gly Ala Leu Ile Tyr Arg Lys Leu Leu Phe Asn Pro  
1 5 10  
aac ggt cct tac cag aaa aag cct gtg cat gaa aaa aaa gaa gtt ttg 248  
Asn Gly Pro Tyr Gln Lys Lys Pro Val His Glu Lys Lys Glu Val Leu  
15 20 25  
tgattttata ttacttttta gtttgatact aagtattaaa catatttctg tattcttcca 308  
aaaaaaaaaaa aaa 321

<210> 116  
<211> 450  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 25..399

<221> sig\_peptide  
<222> 25..186  
<223> Von Heijne matrix  
score 3.5  
seq SILAQVLDQSARA/RL

<400> 116  
ctgctccagc gctgacgccg agcc atg gcg gac gag gag ctt gag gcg ctg 51  
Met Ala Asp Glu Glu Leu Glu Ala Leu  
-50  
agg aga cag agg ctg gcc gag ctg cag gcc aaa cac ggg gat cct ggt 99  
Arg Arg Gln Arg Leu Ala Glu Leu Gln Ala Lys His Gly Asp Pro Gly

```

-45          -40          -35          -30
gat gcg gcc caa cag gaa gca aag cac agg gaa gca gaa atg aga aac      147
Asp Ala Ala Gln Gln Glu Ala Lys His Arg Glu Ala Glu Met Arg Asn
          -25          -20          -15
agt atc tta gcc caa gtt ctg gat cag tcg gcc cgg gcc agg tta agt      195
Ser Ile Leu Ala Gln Val Leu Asp Gln Ser Ala Arg Ala Arg Leu Ser
          -10          -5          1
aac tta gca ctt gta aag cct gaa aaa act aaa gca gta gag aat tac      243
Asn Leu Ala Leu Val Lys Pro Glu Lys Thr Lys Ala Val Glu Asn Tyr
          5          10          15
ctt ata cag atg gca aga tat gga caa cta agt gag aag gta tca gaa      291
Leu Ile Gln Met Ala Arg Tyr Gly Gln Leu Ser Glu Lys Val Ser Glu
          20          25          30          35
caa ggt tta ata gaa atc ctt aaa aaa gta agc caa caa aca gaa aag      339
Gln Gly Leu Ile Glu Ile Leu Lys Lys Val Ser Gln Gln Thr Glu Lys
          40          45          50
aca aca aca gtg aaa ttc aac aga aga aaa gta atg gac tct gat gaa      387
Thr Thr Thr Val Lys Phe Asn Arg Arg Lys Val Met Asp Ser Asp Glu
          55          60          65
gat gac gat tat tgaactacaa gtgctcacag actagaactt aacggaacaa      439
Asp Asp Asp Tyr
          70
gtctaggaca g      450

```

<210> 117  
 <211> 1173  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 10..1137  
 <221> sig\_peptide  
 <222> 10..72  
 <223> Von Heijne matrix  
 score 6.5  
 seq LLTLLPPPPPLYT/RH

<221> polyA\_signal  
 <222> 1144..1149

<221> polyA\_site  
 <222> 1162..1173

```

<400> 117
gagctgctt atg gga cac cgc ttc ctg cgc ggc ctc tta acg ctg ctg ctg      51
          Met Gly His Arg Phe Leu Arg Gly Leu Leu Thr Leu Leu Leu
          -20          -15          -10
ccg ccg cca ccc ctg tat acc cgg cac cgc atg ctc ggt cca gag tcc      99
Pro Pro Pro Pro Leu Tyr Thr Arg His Arg Met Leu Gly Pro Glu Ser
          -5          1          5
gtc ccg ccc cca aaa cga tcc cgc agc aaa ctc atg gca ccg ccc cga      147
Val Pro Pro Pro Lys Arg Ser Arg Ser Lys Leu Met Ala Pro Pro Arg
          10          15          20          25

```

atc ggg acg cac aat ggc acc ttc cac tgc gac gag gca ctg gca tgc	195
Ile Gly Thr His Asn Gly Thr Phe His Cys Asp Glu Ala Leu Ala Cys	
30 35 40	
gca ctg ctt cgc ctc ctg ccg gag tac cgg gat gca gag att gtg cgg	243
Ala Leu Leu Arg Leu Leu Pro Glu Tyr Arg Asp Ala Glu Ile Val Arg	
45 50 55	
acc cgg gat ccc gaa aaa ctc gct tcc tgt gac atc gtg gtg gac gtg	291
Thr Arg Asp Pro Glu Lys Leu Ala Ser Cys Asp Ile Val Val Asp Val	
60 65 70	
ggg ggc gag tac gac cct cgg aga cac cga tat gac cat cac cag agg	339
Gly Gly Glu Tyr Asp Pro Arg Arg His Arg Tyr Asp His His Gln Arg	
75 80 85	
tct ttc aca gag acc atg agc tcc ctg tcc cct ggg agg ccg tgg cag	387
Ser Phe Thr Glu Thr Met Ser Ser Leu Ser Pro Gly Arg Pro Trp Gln	
90 95 100 105	
acc aag ctg agc agt gcg gga ctc atc tat ctg cac ttc ggg cac aag	435
Thr Lys Leu Ser Ser Ala Gly Leu Ile Tyr Leu His Phe Gly His Lys	
110 115 120	
ctg ctg gcc cag ttg ctg ggc act agt gaa gag gac agc atg gtg ggc	483
Leu Leu Ala Gln Leu Leu Gly Thr Ser Glu Glu Asp Ser Met Val Gly	
125 130 135	
acc ctc tat gac aag atg tat gag aac ttt gtg gag gag gtg gat gct	531
Thr Leu Tyr Asp Lys Met Tyr Glu Asn Phe Val Glu Glu Val Asp Ala	
140 145 150	
gtg gac aat ggg atc tcc cag tgg gca gag ggg gag cct cga tat gca	579
Val Asp Asn Gly Ile Ser Gln Trp Ala Glu Gly Glu Pro Arg Tyr Ala	
155 160 165	
ctg acc act acc ctg agt gca cga gtt gct cga ctt aat cct acc tgg	627
Leu Thr Thr Thr Leu Ser Ala Arg Val Ala Arg Leu Asn Pro Thr Trp	
170 175 180 185	
aac cac ccc gac caa gac act gag gca ggg ttc aag cgt gca atg gat	675
Asn His Pro Asp Gln Asp Thr Glu Ala Gly Phe Lys Arg Ala Met Asp	
190 195 200	
ctg gtt caa gag gag ttt ctg cag aga tta gat ttc tac caa cac agc	723
Leu Val Gln Glu Glu Phe Leu Gln Arg Leu Asp Phe Tyr Gln His Ser	
205 210 215	
ggg ctg cca gcc cgg gcc ttg gtg gaa gag gcc ctt gcc cag cga ttc	771
Trp Leu Pro Ala Arg Ala Leu Val Glu Glu Ala Leu Ala Gln Arg Phe	
220 225 230	
cag gtg gac cca agt gga gag att gtg gaa ctg gcg aaa ggt gca tgt	819
Gln Val Asp Pro Ser Gly Glu Ile Val Glu Leu Ala Lys Gly Ala Cys	
235 240 245	
ccc tgg aag gag cat ctc tac cac ctg gaa tct ggg ctg tcc cct cca	867
Pro Trp Lys Glu His Leu Tyr His Leu Glu Ser Gly Leu Ser Pro Pro	
250 255 260 265	
gtg gcc atc ttc ttt gtt atc tac act gac cag gct gga cag tgg cga	915
Val Ala Ile Phe Phe Val Ile Tyr Thr Asp Gln Ala Gly Gln Trp Arg	
270 275 280	
ata cag tgt gtg ccc aag gag ccc cac tca ttc caa agc cgg ctg ccc	963
Ile Gln Cys Val Pro Lys Glu Pro His Ser Phe Gln Ser Arg Leu Pro	
285 290 295	
ctg cca gag cca tgg cgg ggt ctt cgg gag gag gcc ctg gac cag gtc	1011
Leu Pro Glu Pro Trp Arg Gly Leu Arg Asp Glu Ala Leu Asp Gln Val	
300 305 310	
agt ggg atc cct ggc tgc atc ttc gtc cat gca agc ggc ttc att ggc	1059
Ser Gly Ile Pro Gly Cys Ile Phe Val His Ala Ser Gly Phe Ile Gly	
315 320 325	

```
ggt cac cgc acc cga gag ggt gcc ttg agc atg gcc cgt gcc acc ttg 1107
Gly His Arg Thr Arg Glu Gly Ala Leu Ser Met Ala Arg Ala Thr Leu
330          335          340          345
gcc cag cgc tca tac ctc cca caa atc tcc tagtctaata aaaccttoca 1157
Ala Gln Arg Ser Tyr Leu Pro Gln Ile Ser
          350          355
tctcaaaaaa aaaaaa 1173
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<210> 118  
<211> 785  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 72..704

<221> sig\_peptide  
<222> 72..161  
<223> Von Heijne matrix  
score 13.2  
seq LLLLSTLVIPSAA/AP

<221> polyA\_signal  
<222> 772..777

<400> 118

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cggaatccgg gagtccggtg acccgggctg tggcttagca taaaggcggg gccagaaga 60
aggggcgggg t atg gga gaa gcc tcc cca cct gcc ccc gca agg cgg cat 110
Met Gly Glu Ala Ser Pro Pro Ala Pro Ala Arg Arg His
-30          -25          -20
ctg ctg gtc ctg ctg ctg ctc ctc tct acc ctg gtg atc ccc tcc gct 158
Leu Leu Val Leu Leu Leu Leu Leu Ser Thr Leu Val Ile Pro Ser Ala
-15          -10          -5
gca gct cct atc cat gat gct gac gcc caa gag agc tcc ttg ggt ctc 206
Ala Ala Pro Ile His Asp Ala Asp Ala Gln Glu Ser Ser Leu Gly Leu
1          5          10          15
aca ggc ctc cag agc cta ctc caa ggc ttc agc cga ctt ttc ctg aaa 254
Thr Gly Leu Gln Ser Leu Leu Gln Gly Phe Ser Arg Leu Phe Leu Lys
20          25          30
ggt aac ctg ctt cgg ggc ata gac agc tta ttc tct gcc ccc atg gac 302
Gly Asn Leu Leu Arg Gly Ile Asp Ser Leu Phe Ser Ala Pro Met Asp
35          40          45
ttc cgg ggc ctc cct ggg aac tac cac aaa gag gag aac cag gag cac 350
Phe Arg Gly Leu Pro Gly Asn Tyr His Lys Glu Glu Asn Gln Glu His
50          55          60
cag ctg ggg aac aac acc ctc tcc agc cac ctc cag atc gac aag gta 398
Gln Leu Gly Asn Asn Thr Leu Ser Ser His Leu Gln Ile Asp Lys Val
65          70          75
ccc agg atg gag gag aag gag gcc ctg gta ccc atc cag aag gcc acg 446
Pro Arg Met Glu Glu Lys Glu Ala Leu Val Pro Ile Gln Lys Ala Thr
80          85          90          95
gac agc ttc cac aca gaa ctc cat ccc cgg gtg gcc ttc tgg atc att 494
Asp Ser Phe His Thr Glu Leu His Pro Arg Val Ala Phe Trp Ile Ile
100          105          110
```

```

aag ctg cca cgg cgg agg tcc cac cag gat gcc ctg gag ggc ggc cac      542
Lys Leu Pro Arg Arg Arg Ser His Gln Asp Ala Leu Glu Gly Gly His
      115                      120                      125
tgg ctc agc gag aag cga cac cgc ctg cag gcc atc cgg gat gga ctc      590
Trp Leu Ser Glu Lys Arg His Arg Leu Gln Ala Ile Arg Asp Gly Leu
      130                      135                      140
cgc aag ggg acc cac aag gac gtc cta gaa gag ggg acc gag agc tcc      638
Arg Lys Gly Thr His Lys Asp Val Leu Glu Glu Gly Thr Glu Ser Ser
      145                      150                      155
tcc cac tcc agg ctg tcc ccc cga aag acc cac tta ctg tac atc ctc      686
Ser His Ser Arg Leu Ser Pro Arg Lys Thr His Leu Leu Tyr Ile Leu
      160                      165                      170                      175
agg ccc tct cgg cag ctg taggggtggg gaccggggag cacctgcctg      734
Arg Pro Ser Arg Gln Leu
      180
tagcccccac cagaccctgc cccaagcacc atatggaaat aaagttcttt c      785

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<210> 119

<211> 559

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 44..505

<221> sig\_peptide

<222> 44..223

<223> Von Heijne matrix

score 4

seq LVRRTLLVAALRA/WM

<400> 119

```

agcaaccaga gggagatgat cacctgaacc actgctccaa acc atg ggc agt aaa      55
                                     Met Gly Ser Lys
                                     -60
tgc tgt aaa ggt ggt cca gat gaa gat gca gta gaa aga cag agg cgg      103
Cys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu Arg Gln Arg Arg
-55                      -50                      -45
cag aag ttg ctt ctt gca caa ctg cat cac aga aaa agg gtg aag gca      151
Gln Lys Leu Leu Leu Ala Gln Leu His His Arg Lys Arg Val Lys Ala
-40                      -35                      -30                      -25
gct ggg cag atc cag gcc tgg tgg cgt ggg gtc ctg gtg cgc agg acc      199
Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu Val Arg Arg Thr
-20                      -15                      -10
ctg ctg gtt gct gcc ctc agg gcc tgg atg att cag tgc tgg tgg agg      247
Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln Cys Trp Trp Arg
-5                      1                      5
acg ttg gtg cag aga cgg atc cgt cag cgg cgg cag gcc ctg ttg agg      295
Thr Leu Val Gln Arg Arg Ile Arg Gln Arg Arg Gln Ala Leu Leu Arg
10                      15                      20
gtc tac gtc atc cag gag cag gcg acg gtc aag ctc cag tcc tgc atc      343
Val Tyr Val Ile Gln Glu Gln Ala Thr Val Lys Leu Gln Ser Cys Ile
25                      30                      35                      40
cgc atg tgg cag tgc cgg caa tgt tac cgc caa atg tgc aat gct ctc      391

```

```

Arg Met Trp Gln Cys Arg Gln Cys Tyr Arg Gln Met Cys Asn Ala Leu
      45              50              55
tgc ttg ttc cag gtc cca gag agc agc ctt gcc ttc cag act gat ggc      439
Cys Leu Phe Gln Val Pro Glu Ser Ser Leu Ala Phe Gln Thr Asp Gly
      60              65              70
ttt tta cag gtc caa tat gca atc cct tca aag cag cca gag ttc cac      487
Phe Leu Gln Val Gln Tyr Ala Ile Pro Ser Lys Gln Pro Glu Phe His
      75              80              85
att gaa atc cta tca atc tgaaaggcct ggggcatgga gaacaggctg      535
Ile Glu Ile Leu Ser Ile
      90
cactacccta ataaatgtct gacc      559

```

<210> 120  
 <211> 770  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 25..393  
 <221> sig\_peptide  
 <222> 25..150  
 <223> Von Heijne matrix  
       score 4.6  
       seq LDPAVSLSAPAFA/SA  
 <221> polyA\_signal  
 <222> 734..739  
 <221> polyA\_site  
 <222> 757..770

```

<400> 120
CGcagaaagg agagacacac atac atg aaa gga gga gct ttc tcc aat ctt      51
      Met Lys Gly Gly Ala Phe Ser Asn Leu
      -40              -35
aat gat tcc cag ctc tca gcc tcg ttt ctg caa ccc agc ctg caa gca      99
Asn Asp Ser Gln Leu Ser Ala Ser Phe Leu Gln Pro Ser Leu Gln Ala
      -30              -25              -20
aac tgt cct gct ttg gac cct gct gtg tca ctc tcc gca cca gcc ttt      147
Asn Cys Pro Ala Leu Asp Pro Ala Val Ser Leu Ser Ala Pro Ala Phe
      -15              -10              -5
gcc tct gct ctt cgc tct atg aag tcc tcc cag gct gca cgg aag gac      195
Ala Ser Ala Leu Arg Ser Met Lys Ser Ser Gln Ala Ala Arg Lys Asp
      1              5              10              15
gac ttt ctc agg tct ctt agt gat gga gac tca ggg aca tca gaa cac      243
Asp Phe Leu Arg Ser Leu Ser Asp Gly Asp Ser Gly Thr Ser Glu His
      20              25              30
atc tca gcg gtg gtg act agc cct cgg att tcc tgc cat ggt gct gcc      291
Ile Ser Ala Val Val Thr Ser Pro Arg Ile Ser Cys His Gly Ala Ala
      35              40              45
att ccc acc gcc cgt gcc ctc tgc cta ggc tgt tcc tgc tgc acc gaa      339
Ile Pro Thr Ala Arg Ala Leu Cys Leu Gly Cys Ser Cys Cys Thr Glu

```

```

      50              55              60
cgc ctc ctc ctg cca ccg ccc tcc ctc ctt tct tta gaa gcc cct gcc      387
Arg Leu Leu Leu Pro Pro Pro Ser Leu Leu Ser Leu Glu Ala Pro Ala
      65              70              75
agc acc tgagctctct gctgattgct gttcctccca gtctgtggaa gctttgcccc      443
Ser Thr
      80
tatgctttcc ttaaaagggt tctgggcagg gcaggcgccc ccattttotca gggatccccc      503
ccaggacaac gccttttcct tgtgtcttca gctctcctta ccagatatct atatatttgt      563
atatattcag tttcaccaac aatgcatcaa gtactttttt ttttaagtaa agaaccgcag      623
tcatcgaact ggagccccc tgaattccctc cccctcgcc ccccaaactct ggcacctgcc      683
caaggatatcc tcagaaccat ttgggggtgct ctttggcatt ggataataga aataaaattt      743
tacctctttc tacaaaaaaa aaaaaaac      770
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<210> 121  
<211> 1213  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 58..1095  
  
<221> sig\_peptide  
<222> 58..114  
<223> Von Heijne matrix  
score 5.4  
seq LSHLLPSLRQVIQ/EP

<221> polyA\_site  
<222> 1202..1213

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<400> 121
cctggcctttg cctttgccct gctgtgtgat ctagctccc tgcccaggcc cacagcc      57
atg gcc atg gcc cag aaa ctc agc cac ctc ctg ccg agt ctg cgg cag      105
Met Ala Met Ala Gln Lys Leu Ser His Leu Leu Pro Ser Leu Arg Gln
      -15              -10              -5
gtc atc cag gag cct cag cta tct ctg cag cca gag cct gtc ttc acg      153
Val Ile Gln Glu Pro Gln Leu Ser Leu Gln Pro Glu Pro Val Phe Thr
      1              5              10
gtg gat cga gct gag gtg ccg ccg ctc ttc tgg aag ccg tac atc tat      201
Val Asp Arg Ala Glu Val Pro Pro Leu Phe Trp Lys Pro Tyr Ile Tyr
      15              20              25
gcg gcc tac cgg ccg ctg cat cag acc tgg cgc ttc tat ttc cgc acg      249
Ala Gly Tyr Arg Pro Leu His Gln Thr Trp Arg Phe Tyr Phe Arg Thr
      30              35              40              45
ctg ttc cag cag cac aac gag gcc gtg aat gtc tgg acc cac ctg ctg      297
Leu Phe Gln Gln His Asn Glu Ala Val Asn Val Trp Thr His Leu Leu
      50              55              60
gcg gcc ctg gta ctg ctg ctg ccg ctg gcc ctc ttt gtg gag acc gtg      345
Ala Ala Leu Val Leu Leu Leu Arg Leu Ala Leu Phe Val Glu Thr Val
      65              70              75
gac ttc tgg gga gac cca cac gcc ctg ccc ctc ttc atc att gtc ctt      393
Asp Phe Trp Gly Asp Pro His Ala Leu Pro Leu Phe Ile Ile Val Leu
      80              85              90
```



```

gcc tct ttc acc tac ctc tcc ctc agt gcc ttg gct cac ctc ctg cag      441
Ala Ser Phe Thr Tyr Leu Ser Leu Ser Ala Leu Ala His Leu Leu Gln
  95                               100                               105

gcc aag tct gag ttc tgg cat tac agc ttc ttc ttc ctg gac tat gtg      489
Ala Lys Ser Glu Phe Trp His Tyr Ser Phe Phe Phe Leu Asp Tyr Val
110                               115                               120                               125

ggg gtg gcc gtg tac cag ttt ggc agt gcc ttg gca cac ttc tac tat      537
Gly Val Ala Val Tyr Gln Phe Gly Ser Ala Leu Ala His Phe Tyr Tyr
                               130                               135                               140

gct atc gag ccc gcc tgg cat gcc cag gtg cag gct gtt ttt ctg ccc      585
Ala Ile Glu Pro Ala Trp His Ala Gln Val Gln Ala Val Phe Leu Pro
                               145                               150                               155

atg gct gcc ttt ctc gcc tgg ctt tcc tgc att ggc tcc tgc tat aac      633
Met Ala Ala Phe Leu Ala Trp Leu Ser Cys Ile Gly Ser Cys Tyr Asn
160                               165                               170

aag tac atc cag aaa cca ggc ctg ctg ggc cgc aca tgc cag gag gtg      681
Lys Tyr Ile Gln Lys Pro Gly Leu Leu Gly Arg Thr Cys Gln Glu Val
175                               180                               185

ccc tcc gtc ctg gcc tac gca ctg gac att agt cct gtg gtg cat cgt      729
Pro Ser Val Leu Ala Tyr Ala Leu Asp Ile Ser Pro Val Val His Arg
190                               195                               200                               205

atc ttc gtg tcc tcc gac ccc acc acg gat gat cca gct ctt ctc tac      777
Ile Phe Val Ser Ser Asp Pro Thr Thr Asp Asp Pro Ala Leu Leu Tyr
                               210                               215                               220

cac aag tgc cag gtg gtc ttc ttt ctg ctg gct gct gcc ttc ttc tct      825
His Lys Cys Gln Val Val Phe Phe Leu Leu Ala Ala Ala Phe Phe Ser
                               225                               230                               235

acc ttc atg ccc gag cgc tgg ttc cct ggc agc tgc cat gtc ttc ggg      873
Thr Phe Met Pro Glu Arg Trp Phe Pro Gly Ser Cys His Val Phe Gly
240                               245                               250

cag ggc cac caa ctt ttc cat atc ttc ttg gtg ctg tgc acg ctg gct      921
Gln Gly His Gln Leu Phe His Ile Phe Leu Val Leu Cys Thr Leu Ala
255                               260                               265

cag ctg gag gct gtg gca ctg gac tat gag gcc cga cgg ccc atc tat      969
Gln Leu Glu Ala Val Ala Leu Asp Tyr Glu Ala Arg Arg Pro Ile Tyr
270                               275                               280                               285

gag cct ctg cac acg cac tgg cct cac aac ttt tct ggc ctc ttc ctg      1017
Glu Pro Leu His Thr His Trp Pro His Asn Phe Ser Gly Leu Phe Leu
290                               295                               300

ctc acg gtg ggc agc agc atc ctc act gca ttc ctc ctg agc cag ctg      1065
Leu Thr Val Gly Ser Ser Ile Leu Thr Ala Phe Leu Leu Ser Gln Leu
305                               310                               315

gta cag cgc aaa ctt gat cag aag acc aag tgaaggggga tggcatctgg      1115
Val Gln Arg Lys Leu Asp Gln Lys Thr Lys
320                               325

tagggagggga ggtatagttg ggggacaggg gtctgggttt ggctccaagt gggaacaagg      1175
cctggtaaaag ttgtttgtgt ctggccaaaa aaaaaaaaaa
1213

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<210> 122

<211> 1318

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 31..660

<221> sig\_peptide

<222> 31..90

<223> Von Heijne matrix

score 5.4

seq AFVIACVLSLIST/IY

<221> polyA\_signal

<222> 1288..1293

<221> polyA\_site

<222> 1307..1318

<400> 122

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ggaggatggg cgagcagtct gaatgccaga atg gat aac cgt ttt gct aca gca      54
                               Met Asp Asn Arg Phe Ala Thr Ala
                               -20                               -15
ttt gta att gct tgt gtg ctt agc ctc att tcc acc atc tac atg gca      102
Phe Val Ile Ala Cys Val Leu Ser Leu Ile Ser Thr Ile Tyr Met Ala
      -10                               -5                               1
gct tcc att ggc aca gac ttc tgg tat gag tat cga agt cca gtt caa      150
Ala Ser Ile Gly Thr Asp Phe Trp Tyr Glu Tyr Arg Ser Pro Val Gln
      5                               10                               15                               20
gaa aat tcc agt gat ttg aat aaa agc atc tgg gat gaa ttc att agt      198
Glu Asn Ser Ser Asp Leu Asn Lys Ser Ile Trp Asp Glu Phe Ile Ser
      25                               30                               35
gat gag gca gat gaa aag act tat aat gat gca ctt ttt cga tac aat      246
Asp Glu Ala Asp Glu Lys Thr Tyr Asn Asp Ala Leu Phe Arg Tyr Asn
      40                               45                               50
ggc aca gtg gga ttg tgg aga cgg tgt atc acc ata ccc aaa aac atg      294
Gly Thr Val Gly Leu Trp Arg Arg Cys Ile Thr Ile Pro Lys Asn Met
      55                               60                               65
cat tgg tat agc cca cca gaa agg aca gag tca ttt gat gtg gtc aca      342
His Trp Tyr Ser Pro Pro Glu Arg Thr Glu Ser Phe Asp Val Val Thr
      70                               75                               80
aaa tgt gtg agt ttc aca cta act gag cag ttc atg gag aaa ttt gtt      390
Lys Cys Val Ser Phe Thr Leu Thr Glu Gln Phe Met Glu Lys Phe Val
      85                               90                               95                               100
gat ccc gga aac cac aat agc ggg att gat ctc ctt agg acc tat ctt      438
Asp Pro Gly Asn His Asn Ser Gly Ile Asp Leu Leu Arg Thr Tyr Leu
      105                               110                               115
tgg cgt tgc cag ttc ctt tta cct ttt gtg agt tta ggt ttg atg tgc      486
Trp Arg Cys Gln Phe Leu Leu Pro Phe Val Ser Leu Gly Leu Met Cys
      120                               125                               130
ttt ggg gct ttg atc gga ctt tgt gct tgc att tgc cga agc tta tat      534
Phe Gly Ala Leu Ile Gly Leu Cys Ala Cys Ile Cys Arg Ser Leu Tyr
      135                               140                               145
ccc acc att gcc acg ggc att ctc cat ctc ctt gca gtg aca aag gag      582
Pro Thr Ile Ala Thr Gly Ile Leu His Leu Leu Ala Val Thr Lys Glu
      150                               155                               160
agc atg ctt cca gct gga gct gag tcc aag cac aca gcc act cct gca      630
Ser Met Leu Pro Ala Gly Ala Glu Ser Lys His Thr Ala Thr Pro Ala
      165                               170                               175                               180
cac gca tgc gtg caa aca ggg aag ccc aag taggagaaga ggaaagaggt      680
His Ala Cys Val Gln Thr Gly Lys Pro Lys
      185                               190
```

tgtagggatt	tgggaagaac	cttgattatt	ccctggagga	aaagacaaat	ctacttccct	740
gaaatcacc	tcgaatctac	ttccaccctc	agaacttaaa	atgaactgca	tccttttttt	800
catcttcttt	tcttctccag	tgaatatgat	ctccaaaccc	ttattttttc	tttgaactgt	860
aaaattttcca	ctcatggacg	atgcaaccaa	cagatgcaat	ctctgagaag	atgaaaattg	920
ggacctctta	ttataaaatt	gacctagctg	gactcaggaa	accagggaag	aagtcaatgc	980
aggcatttaa	aatgtaaagt	tttttctggt	taaatctatt	tatttttctt	gtaggttgag	1040
tatttcttcc	cagtttttct	gctctggtgt	ataacaaaca	ggtcaaaatt	tcccatcttt	1100
cctcctgata	gtagttgaat	cctaccttgc	atacttaatg	catagtgaag	tggcatctag	1160
cagaaataca	cacccccaaa	acacaccacc	atttcattag	gtgccccaaa	aattctgtat	1220
ttagcttatt	tattttattgt	tatttttgct	ttttcttaac	ccactatata	ttgactgcaa	1280
acgaattaat	aaattatccc	ttctggaaaa	aaaaaaaa			1318

<210> 123

<211> 853

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 31..582

<221> sig\_peptide

<222> 31..90

<223> Von Heijne matrix

score 5.4

seq AFVIACVLSLIST/IY

<221> polyA\_signal

<222> 816..821

<221> polyA\_site

<222> 840..853

<400> 123

ggaggatggg	cgagcagttct	gaatgccaga	atg	gat	aac	cgt	ttt	gct	aca	gca	54
			Met	Asp	Asn	Arg	Phe	Ala	Thr	Ala	
			-20				-15				
ttt	gta	att	gct	tgt	gtg	ctt	agc	ctc	att	tcc	acc
Phe	Val	Ile	Ala	Cys	Val	Leu	Ser	Leu	Ile	Ser	Thr
			-10			-5				1	
gcc	tcc	att	ggc	aca	gac	ttc	tgg	tat	gaa	tat	cga
Ala	Ser	Ile	Gly	Thr	Asp	Phe	Trp	Tyr	Glu	Tyr	Arg
5			10				15			20	
gaa	aat	tcc	agt	gat	ttg	aat	aaa	agc	atc	tgg	gat
Glu	Asn	Ser	Ser	Asp	Leu	Asn	Lys	Ser	Ile	Trp	Asp
			25				30			35	
gat	gaa	gca	gat	gaa	aag	act	tat	aat	gat	gca	cct
Asp	Glu	Ala	Asp	Glu	Lys	Thr	Tyr	Asn	Asp	Ala	Pro
			40				45			50	
ggc	aca	gtg	gga	ttg	tgg	aga	cgg	tgt	atc	acc	ata
Gly	Thr	Val	Gly	Leu	Trp	Arg	Arg	Cys	Ile	Thr	Ile
			55			60				65	
cat	tgg	tat	agc	cca	cca	gaa	agg	aca	gag	tca	ttt
His	Trp	Tyr	Ser	Pro	Pro	Glu	Arg	Thr	Glu	Ser	Phe
			70			75				80	

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aaa tgt gtg agt ttc aca cta act gag cag ttc atg gag aaa ttt gtt      390
Lys Cys Val Ser Phe Thr Leu Thr Glu Gln Phe Met Glu Lys Phe Val
85          90          95          100
gat ccc gga aac cac aat agc ggg att gat ctc ctt agg acc tat ctt      438
Asp Pro Gly Asn His Asn Ser Gly Ile Asp Leu Leu Arg Thr Tyr Leu
          105          110          115
tgg cgt tgc cag ttc ctt tta cct ttt gtg agt tta ggt ttg atg tgc      486
Trp Arg Cys Gln Phe Leu Leu Pro Phe Val Ser Leu Gly Leu Met Cys
          120          125          130
ttt ggg gct ttg atc gga ctt tgt gct tgc att tgc cga agc tta tat      534
Phe Gly Ala Leu Ile Gly Leu Cys Ala Cys Ile Cys Arg Ser Leu Tyr
          135          140          145
ccc acc att gcc acg ggc att ctc cat ctc ctt gca gat acc atg ctg      582
Pro Thr Ile Ala Thr Gly Ile Leu His Leu Leu Ala Asp Thr Met Leu
          150          155          160
tgaagtccag gccacatgga ggtgtcctgt gtagatgctc cagctgaaat cccaagctaa      642
gctcccaact gacagccaac atcattttcca gccatgtgtg ggagccatcc tggatgtcca      702
gccttaacaa gccttcagag gacttcagcc acagctatta tcttactaca tccttgtgag      762
actctaataa agaaccaact agctgagccc aatcaaccta tggaactgat agaaataaaa      822
tgaattgttg ttttgcgaaa aaaaaaaaaa a                                     853
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<210> 124
<211> 826
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 15..695
<221> sig_peptide
<222> 15..80
<223> Von Heijne matrix
      score 8.5
      seq AALLLGLMMVVTG/DE
<221> polyA_signal
<222> 795..800
<221> polyA_site
<222> 814..826
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<400> 124
aaccagaggt gccc atg ggt tgg aca atg agg ctg gtc aca gca gca ctg      50
          Met Gly Trp Thr Met Arg Leu Val Thr Ala Ala Leu
          -20          -15
tta ctg ggt ctc atg atg gtg gtc act gga gac gag gat gag aac agc      98
Leu Leu Gly Leu Met Val Val Thr Gly Asp Glu Asp Glu Asn Ser
-10          -5          1          5
ccg tgt gcc cat gag gcc ctc ctg gac gag gac acc ctc ttt tgc cag      146
Pro Cys Ala His Glu Ala Leu Leu Asp Glu Asp Thr Leu Phe Cys Gln
          10          15          20
ggc ctt gaa gtt ttc tac cca gag ttg ggg aac att ggc tgc aag gtt      194
Gly Leu Glu Val Phe Tyr Pro Glu Leu Gly Asn Ile Gly Cys Lys Val
          25          30          35
```

```

gtt cct gat tgt aac aac tac aga cag aag atc acc tcc tgg atg gag      242
Val Pro Asp Cys Asn Asn Tyr Arg Gln Lys Ile Thr Ser Trp Met Glu
  40                      45                      50
ccg ata gtc aag ttc ccg ggg gcc gtg gac ggc gca acc tat atc ctg      290
Pro Ile Val Lys Phe Pro Gly Ala Val Asp Gly Ala Thr Tyr Ile Leu
  55                      60                      65                      70
gtg atg gtg gat cca gat gcc cct agc aga gca gaa ccc aga cag aga      338
Val Met Val Asp Pro Asp Ala Pro Ser Arg Ala Glu Pro Arg Gln Arg
  75                      80                      85
ttc tgg aga cat tgg ctg gta aca gat atc aag ggc gcc gac ctg aag      386
Phe Trp Arg His Trp Leu Val Thr Asp Ile Lys Gly Ala Asp Leu Lys
  90                      95                      100
aaa ggg aag att cag ggc cag gag tta tca gcc tac cag gct ccc tcc      434
Lys Gly Lys Ile Gln Gly Gln Glu Leu Ser Ala Tyr Gln Ala Pro Ser
  105                      110                      115
cca ccg gca cac agt ggc ttc cat cgc tac cag ttc ttt gtc tat ctt      482
Pro Pro Gln Ala His Ser Gly Phe His Arg Tyr Gln Phe Phe Val Tyr Leu
  120                      125                      130
cag gaa gga aag gtc atc tct ctc ctt ccc aag gaa aac aaa act cga      530
Gln Glu Gly Lys Val Ile Ser Leu Leu Pro Lys Glu Asn Lys Thr Arg
  135                      140                      145                      150
ggc tct tgg aaa atg gac aga ttt ctg aac cgt ttc cac ctg ggc gaa      578
Gly Ser Trp Lys Met Asp Arg Phe Leu Asn Arg Phe His Leu Gly Glu
  155                      160                      165
cct gaa gca agc acc cag ttc atg acc cag aac tac cag gac tca cca      626
Pro Glu Ala Ser Thr Gln Phe Met Thr Gln Asn Tyr Gln Asp Ser Pro
  170                      175                      180
acc ctc cag gct ccc aga gaa agg gcc agc gag ccc aag cac aaa aac      674
Thr Leu Gln Ala Pro Arg Glu Arg Ala Ser Glu Pro Lys His Lys Asn
  185                      190                      195
cag gcg gag ata gct gcc tgc tagatagccg gctttgccat ccgggcatgt      725
Gln Ala Glu Ile Ala Ala Cys
  200                      205
ggccacactg cccaccacog acgatgtggg tatggaaccc cctctggata cagaaccct      785
Tctttttccaa ataaaaaaaa aatcatccaa aaaaaaaaaa a      826

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<210> 125

<211> 571

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 74..295

<221> sig\_peptide

<222> 74..196

<223> Von Heijne matrix

score 5.4

seq RLLYIGFLGYCSG/LI

<221> polyA\_signal

<222> 545..550

<221> polyA\_site

<222> 561..571

<400> 125

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cgggtagtgg tcgtcgtggg tttccttgta gttcgtgggc tgagaccagg cctcaagtgg      60
aaacggcgctc acc atg atc gca cgg cgg aac cca gta ccc tta cgg ttt      109
          Met Ile Ala Arg Arg Asn Pro Val Pro Leu Arg Phe
          -40          -35          -30
ctg ccg gat gag gcc cgg agc ctg ccc ccg ccc aag ctg acc gac ccg      157
Leu Pro Asp Glu Ala Arg Ser Leu Pro Pro Pro Lys Leu Thr Asp Pro
          -25          -20          -15
cgg ctc ctc tac atc ggc ttc ttg ggc tac tgc tcc ggc ctg att gat      205
Arg Leu Leu Tyr Ile Gly Phe Leu Gly Tyr Cys Ser Gly Leu Ile Asp
          -10          -5          1
aac ctg atc cgg cgg agg ccg atc gcg acg gct ggt ttg cat cgc cag      253
Asn Leu Ile Arg Arg Arg Pro Ile Ala Thr Ala Gly Leu His Arg Gln
          5          10          15
ctt cta tat att acg gcc ttt ttt ttg ctg gat att atc ttg      295
Leu Leu Tyr Ile Thr Ala Phe Phe Leu Leu Asp Ile Ile Leu
          20          25          30
taaaacgtga agactacctg tatgctgtga gggaccgtga aatgttttga tatatgaaat      355
tacatccaga ggattttcct gaagaagata agaaaacata tgggtgaaatt tttgaaaaat      415
tccatccaat acgttgaagt cttcaaaatg cttgctccag tttcactgat acctgctgtt      475
cctgaatttg atggaacatg tttottatga cagttgaagc ttatgctaatt ctgtatgttg      535
acaccttgta attaaaatac gtacccaaaaa aaaaaa      571
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<210> 126

<211> 659

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 440..658

<221> polyA\_signal

<222> 601..606

<400> 126

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cgccctaaga gctgggaggt ggtgcctctc acccagctaa ttgctctcta gcccttggcc      60
ttcacagggtg ttggtgcctg ccgtgaacgc attctgacct gggccgtatc tgtctcccaa      120
gactttgtgc ctatggttgg ggacagagtg aggtcgttgc cttgacgacg acagcatgcg      180
gcccggtggc ctcctaagtg tgagcttgcg gcggaccgag gccacactgc ctccctgcct      240
gcttgcacca ggactcgtga ctgcgtccgc agaagaaatc acaacagcgc tgggaattgct      300
agtttgctag gcagcatctt ttggacctgc gaaccatatg catttcacct caaatctggt      360
tccaagttga aaaccttttg gtctttctat gcgaacggat tgaagaaacg caaaaagttt      420
ctacggactt taaattaaa atg gaa aaa tat gaa aac ctg ggt ttg gtt gga      472
          Met Glu Lys Tyr Glu Asn Leu Gly Leu Val Gly
          1          5          10
gaa ggg agt tat gga atg gtg atg aag tgt agg aat aaa gat act gga      520
Glu Gly Ser Tyr Gly Met Val Met Lys Cys Arg Asn Lys Asp Thr Gly
          15          20          25
aga att gtg gcc ata aag aag ttc tta gaa agt gac gat gac aaa atg      568
Arg Ile Val Ala Ile Lys Lys Phe Leu Glu Ser Asp Asp Asp Lys Met
          30          35          40
gtt aaa aag att gca atg cga gaa gtc aag tta cta aag caa ctt agg      616
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Val Lys Lys Ile Ala Met Arg Glu Val Lys Leu Leu Lys Gln Leu Arg  
45 50 55  
cat gaa aac ttg gtg aat ctc ttg gaa gtg tgt aaa aaa aaa a 659  
His Glu Asn Leu Val Asn Leu Leu Glu Val Cys Lys Lys Lys  
60 65 70

<210> 127  
<211> 301  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 38..283

<221> sig\_peptide  
<222> 38..85  
<223> Von Heijne matrix  
score 4.1  
seq LLPATSLAGPVLS/TL

<221> polyA\_signal  
<222> 257..262

<400> 127  
cacctgaatc ccaggaaccc tcaatgaggt cttcaag atg aag aga ctg ctg cca 55  
Met Lys Arg Leu Leu Pro  
-15  
gct acc agc ctg gct ggc cct gtc ctg tcc acc ctc att gcc cca act 103  
Ala Thr Ser Leu Ala Gly Pro Val Leu Ser Thr Leu Ile Ala Pro Thr  
-10 -5 1 5  
ccc atg ttg ttt tgt gaa gat aaa agc tgg gat ctt ttt ctt ttt ttt 151  
Pro Met Leu Phe Cys Glu Asp Lys Ser Trp Asp Leu Phe Leu Phe Phe  
10 15 20  
aag tct cac aag aca tgg ggc atc tcc aca aat tta agt tcc tgt cca 199  
Lys Ser His Lys Thr Trp Gly Ile Ser Thr Asn Leu Ser Ser Cys Pro  
25 30 35  
ttt gga aat ttg ttt cta tgt gta cag ttt gtc aga gaa aaa caa agt 247  
Phe Gly Asn Leu Phe Leu Cys Val Gln Phe Val Arg Glu Lys Gln Ser  
40 45 50  
ttt tgt atg aat aca gaa tgt gat tta cgc aag aat tgacaaaaaa 293  
Phe Cys Met Asn Thr Glu Cys Asp Leu Arg Lys Asn  
55 60 65  
aaaaaaaaa 301

<210> 128  
<211> 477  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 121..477

<221> sig\_peptide  
 <222> 121..288  
 <223> Von Heijne matrix  
 score 3.5  
 seq SSCADSFVSSSSS/QP

<400> 128  
 cctcggagca ggcggagtaa agggacttga gcgagccagt tgccggatta ttctatttcc 60  
 cctccctctc tcccgccccg tatctctttt cacccttctc ccaccctcgc togcgtagcc 120  
 atg gcg gag ccg tgc gcg gcc act cag tcc cat tcc atc tcc tgc tgc 168  
 Met Ala Glu Pro Ser Ala Ala Thr Gln Ser His Ser Ile Ser Ser Ser  
 -55 -50 -45  
 tcc ttc gga gcc gag ccg tcc gcg ccc ggc ggc ggc ggc agc cca gga 216  
 Ser Phe Gly Ala Glu Pro Ser Ala Pro Gly Gly Gly Gly Ser Pro Gly  
 -40 -35 -30 -25  
 gcc tgc ccc gcc ctg ggg acg aag agc tgc agc tcc tcc tgt gcg gat 264  
 Ala Cys Pro Ala Leu Gly Thr Lys Ser Cys Ser Ser Ser Cys Ala Asp  
 -20 -15 -10  
 tcc ttt gtt tct tcc tct tcc tct cag cct gta tct cta ttt tgc acc 312  
 Ser Phe Val Ser Ser Ser Ser Ser Ser Gln Pro Val Ser Leu Phe Ser Thr  
 -5 1 5  
 tca caa gag gga ttg agc tct ctt tgc tct gat gag cca tct tca gaa 360  
 Ser Gln Glu Gly Leu Ser Ser Leu Cys Ser Asp Glu Pro Ser Ser Glu  
 10 15 20  
 att atg act tct tcc ttt ctt tca tct tct gaa ata cat aac act ggc 408  
 Ile Met Thr Ser Ser Phe Leu Ser Ser Ser Glu Ile His Asn Thr Gly  
 25 30 35 40  
 ctt aca ata cta cat gga gaa aaa agc cat gtg tta ggg agc cag cct 456  
 Leu Thr Ile Leu His Gly Glu Lys Ser His Val Leu Gly Ser Gln Pro  
 45 50 55  
 att tta gcc aaa aaa aaa aaa 477  
 Ile Leu Ala Lys Lys Lys  
 60

<210> 129  
 <211> 323  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 2..163

<221> polyA\_signal  
 <222> 292..297

<221> polyA\_site  
 <222> 310..323

<400> 129  
 a gct ttc gtg tgg gag cca gct atg gtg cgg atc aat gcg ctg aca gca 49  
 Ala Phe Val Trp Glu Pro Ala Met Val Arg Ile Asn Ala Leu Thr Ala  
 1 5 10 15  
 gcc tct gag gct gcg tgc ctg atc gtg tct gta gat gaa acc atc aag 97



```
Ala Ser Glu Ala Ala Cys Leu Ile Val Ser Val Asp Glu Thr Ile Lys
      20              25              30
aac ccc cgc tgc act gtg gat gct ccc aca gca gca ggc cgg ggc cgt      145
Asn Pro Arg Ser Thr Val Asp Ala Pro Thr Ala Ala Gly Arg Gly Arg
      35              40              45
ggg cgt ggc cgc ccc cac tgagaggcac cccacccatc acatggctgg      193
Gly Arg Gly Arg Pro His
      50
ctggctgctg ggtgcactta ccctccttgg cttgggttact tcattttaca aggaaggggt      253
agtaattggc ccactctctt cttactggag gctattttaa taaaatgtaa gacttcaaaa      313
aaaaaaaaaa      323
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<210> 130  
<211> 1392  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 46..675  
<221> sig\_peptide  
<222> 46..87  
<223> Von Heijne matrix  
score 5.3  
seq LTLGLSIFLAGL/IV  
<221> polyA\_signal  
<222> 1364..1369  
<221> polyA\_site  
<222> 1383..1392

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<400> 130
ctccgagttg ccaccagga aaaagagggc tcctctggga gatgt atg ctt act ctc      57
      Met Leu Thr Leu
tta ggc ctt tca ttc atc ttg gca gga ctt att gtt ggt gga gcc tgc      105
Leu Gly Leu Ser Phe Ile Leu Ala Gly Leu Ile Val Gly Gly Ala Cys
-10              -5              1              5
att tac aag tac ttc atg ccc aag agc acc att tac cgt gga gag atg      153
Ile Tyr Lys Tyr Phe Met Pro Lys Ser Thr Ile Tyr Arg Gly Glu Met
      10              15              20
tgc ttt ttt gat tct gag gat cct gca aat tcc ctt cgt gga gga gag      201
Cys Phe Phe Asp Ser Glu Asp Pro Ala Asn Ser Leu Arg Gly Gly Glu
      25              30              35
cct aac ttc ctg cct gtg act gag gag gct gac att cgt gag gat gac      249
Pro Asn Phe Leu Pro Val Thr Glu Glu Ala Asp Ile Arg Glu Asp Asp
      40              45              50
aac att gca atc att gat gtg cct gtc ccc agt ttc tct gat agt gac      297
Asn Ile Ala Ile Ile Asp Val Pro Val Pro Ser Phe Ser Asp Ser Asp
      55              60              65              70
cct gca gca att att cat gac ttt gaa aag gga atg act gct tac ctg      345
Pro Ala Ala Ile Ile His Asp Phe Glu Lys Gly Met Thr Ala Tyr Leu
      75              80              85
gac ttg ttg ctg ggg atc tgc tat ctg atg ccc ctc aat act tct att      393
```

```

Asp Leu Leu Leu Gly Ile Cys Tyr Leu Met Pro Leu Asn Thr Ser Ile
    90                      95                      100
gtt atg cct cca aaa aat ctg gta gag ctc ttt ggc aaa ctg gcg agt    441
Val Met Pro Pro Lys Asn Leu Val Glu Leu Phe Gly Lys Leu Ala Ser
    105                      110                      115
ggc aga tat ctg cct caa act tat gtg gtt cga gaa gac cta gtt gct    489
Gly Arg Tyr Leu Pro Gln Thr Tyr Val Val Arg Glu Asp Leu Val Ala
    120                      125                      130
gtg gag gaa att cgt gat gtt agt aac ctt ggc atc ttt att tac caa    537
Val Glu Glu Ile Arg Asp Val Ser Asn Leu Gly Ile Phe Ile Tyr Gln
    135                      140                      145                      150
ctt tgc aat aac aga aag tcc ttc cgc ctt cgt cgc aga gac ctc ttg    585
Leu Cys Asn Asn Arg Lys Ser Phe Arg Leu Arg Arg Arg Asp Leu Leu
    155                      160                      165
ctg ggt ttc aac aaa cgt gcc att gat aaa tgc tgg aag att aga cac    633
Leu Gly Phe Asn Lys Arg Ala Ile Asp Lys Cys Trp Lys Ile Arg His
    170                      175                      180
ttc ccc aac gaa ttt att gtt gag acc aag atc tgt caa gag    675
Phe Pro Asn Glu Phe Ile Val Glu Thr Lys Ile Cys Gln Glu
    185                      190                      195
taagaggcaa cagatagagt gtccttggtg ataagaagtc agagatttac aatatgactt    735
taacattaag gtttatggga tactcaagat atttactcat gcatttactc tattgcttat    795
gctttaaaaa aaggaaaaaaa aaaaaactac taaccactgc aagctcttgt caaatttttag    855
ttttaattggc attgcttggt ttttgaaact gaaattacat gagtttcatt tttcctttgc    915
atztataggg tttagatttc tgaaagcagc atgaatatat cacctaacat cctgacaata    975
aattccatcc gttgtttttt ttgtttgttt gttttttctt ttcctttaag taagctcttt 1035
attcatctta tgggtggagca attttaaaat ttgaaatatt ttaaattggt tttgaacttt 1095
ttgtgtaaaa tatatcagat ctcaacattg ttggtttctt ttgtttttca ttttgtacaa 1155
ctttcttgaa tttagaaatt acatctttgc agttctgtta ggtgctctgt aattaacctg 1215
acttatatgt gaacaatttt catgagacag tcatttttta ctaatgcagt gattctttct 1275
cactactatc tgtattgtgg aatgcacaaa attgtgtagg tgctgaatgc tgtaaggagt 1335
ttaggttgta tgaattctac aaccctataa taaattttac tctatacaaa aaaaaaa 1392

<210> 131
<211> 999
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 62..385

<221> polyA_signal
<222> 974..979

<221> polyA_site
<222> 987..999

<400> 131
cctgaatgac ttgaatgttt cccgcctga gctaacagtc catgtgggtg attcagctct    60
g atg gga tgt gtt ttc cag agc aca gaa gac aaa tgt ata ttc aag ata    109
Met Gly Cys Val Phe Gln Ser Thr Glu Asp Lys Cys Ile Phe Lys Ile
    1                      5                      10                      15
gac tgg act ctg tca cca gga gag cac gcc aag gac gaa tat gtg cta    157
Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu

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	20		25		30	
tac tat tac tcc aat ctc agt gtg cct att ggg cgc ttc cag aac cgc						205
Tyr Tyr Tyr Ser Asn Leu Ser Val Pro Ile Gly Arg Phe Gln Asn Arg						
	35		40		45	
gta cac ttg atg ggg gac atc tta tgc aat gat ggc tct ctc ctg ctc						253
Val His Leu Met Gly Asp Ile Leu Cys Asn Asp Gly Ser Leu Leu Leu						
	50		55		60	
caa gat gtg caa gag gct gac cag gga acc tat atc tgt gaa atc cgc						301
Gln Asp Val Gln Glu Ala Asp Gln Gly Thr Tyr Ile Cys Glu Ile Arg						
	65		70		75	80
ctc aaa ggg gag agc cag gtg ttc aag aag gcg gtg gta ctg cat gtg						349
Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val						
	85		90		95	
ctt cca gag gag ccc aaa ggt acg caa atg ctt act taaagagggg						395
Leu Pro Glu Glu Pro Lys Gly Thr Gln Met Leu Thr						
	100		105			
ccaaggggca agagctttca tgtgcaagag gcaaggaaac tgattatctt gagtaaatagc						455
cagccttttg gctaagtact taccacagag tgaatcttca aaaaatgatac ataattatctt						515
cagtcaataa aaatagagtt attttattaa ataaaatatt gataattatt gtattattac						575
tttaaacaca cttccccctc acaaaagccc tgtgaaggat gttttgttca catatatgtc						635
caaatatgtt ttggacacat atttattaaa tggaataaat agtacttgaa ccctggcacc						695
tctgacaaca aagtccatgt tctttttact atgccctaatac acctttcatc agttatccac						755
attgatgcta catctgtatt ttataggtac cctatgttag gtgttctggg ggatagaaaa						815
gaaataagca ggccaggctc agtggctcat gcctgtaatac ctagcatttt gggaggctga						875
ggcagcagaa ctgcctgagc cccagggttc aagactgcag tgagctatga tggcaccact						935
gcattctagc ctgggtgaca gagcaagact ctgtctaaaa taaaaaaaga gaaaaaaaaa						995
aaaa						999

<210> 132

<211> 725

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 422..550

<221> sig\_peptide

<222> 422..475

<223> Von Heijne matrix

score 4.5

seq LRWLMPVIPALWG/AE

<221> polyA\_site

<222> 714..725

<400> 132

tctgagaggg tgggagagaa aattaggggg agaaaggaca gagagagcaa ctaccatcca	60
tagccagata ggtgagtaaa tatatttgca gtaacctatt tgctattcct tgcgtcaact	120
gtgttttaag ttccttccag aatcagagag agtattgcca tccaagaaat cgtttttaga	180
tatgacattt gagctatcat cttgagacca atacctaaaa caatttcagt ttaagaaatg	240
tctaggtatg gtgaaaacac agtttaaaac cagcaaaaca gaatttattg ccctcagcga	300
ataccacaaa tgtacatata ccttgtattt ctgaaagcaa agcaagcatg ccaagtagtt	360
tttattttacc tgtacctata atacagcaag gtgaaacagg atatattttt gaagtttaaa	420
a atg tct tca ggc cgg ctg cgg tgg ctc atg cct gta atc cca gca ctt	469

```

Met Ser Ser Gly Arg Leu Arg Trp Leu Met Pro Val Ile Pro Ala Leu
      -15          -10          -5
tgg gga gcc gag aag ggt gaa tca cct gag gtc agc agt ttt gag acc    517
Trp Gly Ala Glu Lys Gly Glu Ser Pro Glu Val Ser Ser Phe Glu Thr
      1          5          10
agg ctg gcc aac atg gcg aaa ccc tgt ctc tac tgaaaataca aaaattagct    570
Arg Leu Ala Asn Met Ala Lys Pro Cys Leu Tyr
      15          20          25
gggtgtggtg gcgggcgctt gtagtcccag ctacttggga gactgaggca ggagaattgc    630
ttgaacacgg aaggcggaag ttgcagtaag ctgagatcgt gccaccgcac accagcttgg    690
gcaacagagt gagactccct ctcaaaaaaa aaaaaa                                725

```

<210> 133  
 <211> 400  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 124..231

<221> polyA\_site  
 <222> 387..400

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<400> 133
ctcgcctctc ctggcttctg gtatgcacca gcaattcctg gcgttccttg gctcctagaa    60
gcatcactcc tatcacatgg tcattcttcac cctgtgtgtc ttcacactac cttttctctg    120
tgc atg tct gcc cga atc cct ttt tat aag gac acc agt cag att aga        168
Met Ser Ala Arg Ile Pro Phe Tyr Lys Asp Thr Ser Gln Ile Arg
      1          5          10          15
tta ggg tct acc ata ata cct cat ttt aac tta atc acc ttt gta aag        216
Leu Gly Ser Thr Ile Ile Pro His Phe Asn Leu Ile Thr Phe Val Lys
      20          25          30
acc ttt ttc caa ata tagtcactct ctgaggtact gatggtagg atctcaacat        271
Thr Phe Phe Gln Ile
      35
accttttttg ggaggacaca attgaacca taacagggtg tttgcaagga agagttaaaa    331
tttgaaagaa aggtggtatt tgcttagata gatagggcac agctttctag gtgacaaaaa    391
aaaaaaaaa                                400

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<210> 134  
 <211> 1053  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 131..1051

<221> sig\_peptide  
 <222> 131..169  
 <223> Von Heijne matrix  
 score 4.2

seq MLAVSLTVPLLGA/MM

<221> polyA\_signal

<222> 1019..1024

<400> 134

gagcgaggcg	gacgggctgc	gacagcgccg	gcccctgcgg	ccgcaggctcg	tcacagacga	60
tgatggccag	gccccggagg	ctaaggacgg	cagctccttt	agcggcagag	ttttccgagt	120
gaccttcttg	atg ctg gct gtt tct ctc acc gtt ccc ctg ctt gga gcc	169				
	Met Leu Ala Val Ser Leu Thr Val Pro Leu Leu Gly Ala					
	-10	-5				
atg atg ctg ctg gaa tct cct ata gat cca cag cct ctc agc ttc aaa	217					
Met Met Leu Leu Glu Ser Pro Ile Asp Pro Gln Pro Leu Ser Phe Lys						
1	5	10	15			
gaa ccc ccg ctc ttg ctt ggt gtt ctg cat cca aat acg aag ctg cga	265					
Glu Pro Pro Leu Leu Leu Gly Val Leu His Pro Asn Thr Lys Leu Arg						
	20	25	30			
cag gca gaa agg ctg ttt gaa aat caa ctt gtt gga ccg gag tcc ata	313					
Gln Ala Glu Arg Leu Phe Glu Asn Gln Leu Val Gly Pro Glu Ser Ile						
	35	40	45			
gca cat att ggg gat gtg atg ttt act ggg aca gca gat ggc cgg gtc	361					
Ala His Ile Gly Asp Val Met Phe Thr Gly Thr Ala Asp Gly Arg Val						
	50	55	60			
gta aaa ctt gaa aat ggt gaa ata gag acc att gcc cgg ttt ggt tcg	409					
Val Lys Leu Glu Asn Gly Glu Ile Glu Thr Ile Ala Arg Phe Gly Ser						
	65	70	75	80		
ggc cct tgc aaa acc cga gat gat gag cct gtg tgt ggg aga ccc ctg	457					
Gly Pro Cys Lys Thr Arg Asp Asp Glu Pro Val Cys Gly Arg Pro Leu						
	85	90	95			
ggg atc cgt gca ggg ccc aat ggg act ctc ttt gtg gcc gat gca tgc	505					
Gly Ile Arg Ala Gly Pro Asn Gly Thr Leu Phe Val Ala Asp Ala Cys						
	100	105	110			
aag gga cta ttt gaa gta aat ccc tgg aaa cgt gaa gtg aaa ctg ctg	553					
Lys Gly Leu Phe Glu Val Asn Pro Trp Lys Arg Glu Val Lys Leu Leu						
	115	120	125			
ctg tcc tcc gag aca ccc att gag ggg aag aac atg tcc ttt gtg aat	601					
Leu Ser Ser Glu Thr Pro Ile Glu Gly Lys Asn Met Ser Phe Val Asn						
	130	135	140			
gat ctt aca gtc tct cag gat ggg agg aag att tat ttc acc gat tct	649					
Asp Leu Thr Val Ser Gln Asp Gly Arg Lys Ile Tyr Phe Thr Asp Ser						
	145	150	155	160		
agc agc aaa tgg caa aga cga gac tac ctg ctt ctg gtg atg gag ggc	697					
Ser Ser Lys Trp Gln Arg Arg Asp Tyr Leu Leu Leu Val Met Glu Gly						
	165	170	175			
aca gat gac ggg cgc ctg ctg gag tat gat act gtg acc agg gaa gta	745					
Thr Asp Asp Gly Arg Leu Leu Glu Tyr Asp Thr Val Thr Arg Glu Val						
	180	185	190			
aaa gtt tta ttg gac cag ctg cgg ttc ccg aat gga gtc cag ctg tct	793					
Lys Val Leu Leu Asp Gln Leu Arg Phe Pro Asn Gly Val Gln Leu Ser						
	195	200	205			
cct gca gaa gac ttt gtc ctg gtg gca gaa aca acc atg gcc agg ata	841					
Pro Ala Glu Asp Phe Val Leu Val Ala Glu Thr Thr Met Ala Arg Ile						
	210	215	220			
cga aga gtc tac gtt tct ggc ctg atg aag ggc gct gat ctg ttt	889					
Arg Arg Val Tyr Val Ser Gly Leu Met Lys Gly Gly Ala Asp Leu Phe						
	225	230	235	240		
gtg gag aac atg cct gga ttt cca gac aac atc cgg ccc agc agc tct	937					

```
Val Glu Asn Met Pro Gly Phe Pro Asp Asn Ile Arg Pro Ser Ser Ser
      245                250                255
ggg ggg tac tgg gtg ggc atg tcg acc atc cgc cct aac cct ggg ttt      985
Gly Gly Tyr Trp Val Gly Met Ser Thr Ile Arg Pro Asn Pro Gly Phe
      260                265                270
tcc atg ctg gat ttc tta tct gag aga ccc tgg att aaa agg atg att      1033
Ser Met Leu Asp Phe Leu Ser Glu Arg Pro Trp Ile Lys Arg Met Ile
      275                280                285
ttt aag gca aaa aaa aaa aa      1053
Phe Lys Ala Lys Lys Lys
      290
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<210> 135

<211> 1128

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 86..403

<221> sig\_peptide

<222> 86..181

<223> Von Heijne matrix

score 8.8

seq VPMLLLIVGGSFG/LR

<221> polyA\_signal

<222> 1097..1102

<221> polyA\_site

<222> 1117..1128

<400> 135

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cgctcttggtg agagcgtgag ctgctgagat ttgggagtct gcgctaggcc cgcttggagt      60
ctgagccga tggaagagtt cactc atg ttt gca ccc gcg gtg atg cgt gct      112
      Met Phe Ala Pro Ala Val Met Arg Ala
      -30                -25
ttt cgc aag aac aag act ctc ggc tat gga gtc ccc atg ttg ttg ctg      160
Phe Arg Lys Asn Lys Thr Leu Gly Tyr Gly Val Pro Met Leu Leu Leu
      -20                -15                -10
att gtt gga ggt tct ttt ggt ctt cgt gag ttt tct caa atc cga tat      208
Ile Val Gly Gly Ser Phe Gly Leu Arg Glu Phe Ser Gln Ile Arg Tyr
      -5                1                5
gat gct gtg aag agt aaa atg gat cct gag ctt gaa aaa aaa ctg aaa      256
Asp Ala Val Lys Ser Lys Met Asp Pro Glu Leu Glu Lys Lys Leu Lys
      10                15                20                25
gag aat aaa ata tct tta gag tcg gaa tat gag aaa atc aaa gac tcc      304
Glu Asn Lys Ile Ser Leu Glu Ser Glu Tyr Glu Lys Ile Lys Asp Ser
      30                35                40
aag ttt gat gac tgg aag aat att cga gga ccc agg cct tgg gaa gat      352
Lys Phe Asp Asp Trp Lys Asn Ile Arg Gly Pro Arg Pro Trp Glu Asp
      45                50                55
cct gac ctc ctc caa gga aga aat cca gaa agc ctt aag act aag aca      400
Pro Asp Leu Leu Gln Gly Arg Asn Pro Glu Ser Leu Lys Thr Lys Thr
```

60	65	70	
act tgactctgct gattcttttt tccnnntttt ttttttttta aataaaaata			453
Thr			
ctattaactg gacttcctaa tatatacttc tatcaagtgg aaaggaaatt ccaggcccat			513
ggaaacttgg atatgggtaa tttgatgaca aataatcttc actaaaggtc atgtacaggt			573
ttttatactt cccagctatt ccatctgtgg atgaaagtaa caatgttggc cacgtatatt			633
ttacacctcg aaataaaaaa tgtgaatact gctccaaaaa aaaaaaccag taccgtgtag			693
tctctctcgt ggcttggatt tacactgggc aacgtgggtg gaatgtatct ggctcagaac			753
tatgatatac caaacctggc taaaaaactt gaagaaatta aaaaggactt ggatgccaag			813
aagaaacccc ctagtgcatt agactgcctc cagcactgcc ttcaggatat accgattcta			873
ctgctcttga gggcctcgtt tactatctga accaaaagct tttgttttcg tctccagcct			933
cagcacttct cttctttgct agaccctgtg ttttttgctt taaagcaagc aaaatggggc			993
cccaatttga gaactacccg acgtttccaa catactcacc tcttcccata atccctttcc			1053
aactgcatgg gaggttctaa gactggaatt atgggtgctag attagtaaac atgactttta			1113
acgaaaaaaa aaaaa			1128

<210> 136

<211> 254

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 37..162

<221> sig\_peptide

<222> 37..93

<223> Von Heijne matrix

score 9.5

seq LMCLSLCTAFALS/KP

<221> polyA\_signal

<222> 224..229

<221> polyA\_site

<222> 243..254

<400> 136

tggtgtgtgg gggctacgag gaaagatcta attatc atg gac ctg cga cag ttt	54
Met Asp Leu Arg Gln Phe	
	-15
ctt atg tgc ctg tcc ctg tgc aca gcc ttt gcc ttg agc aaa ccc aca	102
Leu Met Cys Leu Ser Leu Cys Thr Ala Phe Ala Leu Ser Lys Pro Thr	
	-10 -5 1
gaa aag aag gac cgt gta cat cat gag cct cag ctc agt gac aag gtt	150
Glu Lys Lys Asp Arg Val His His Glu Pro Gln Leu Ser Asp Lys Val	
	5 10 15
cac aat gat att tgatagaacc aattgttgta cataaaacag atctgcgcat	202
His Asn Asp Ile	
20	
atatatatat gtataaaaaa taataaaata atggaagatg aaaaaaaaaa aa	254

<210> 137

<211> 886  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 31..381

<221> sig\_peptide  
<222> 31..90  
<223> Von Heijne matrix  
score 5.4  
seq AFVIACVLSLIST/IY

<221> polyA\_site  
<222> 875..886

<400> 137

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ggagggatggg cgagcagtcct gaatggcaga atg gat aac cgt ttt gct aca gca      54
          Met Asp Asn Arg Phe Ala Thr Ala
                    -20                      -15
ttt gta att gct tgt gtg ctt agc ctc att tcc acc atc tac atg gca      102
Phe Val Ile Ala Cys Val Leu Ser Leu Ile Ser Thr Ile Tyr Met Ala
          -10                      -5                      1
gcc tcc att ggc aca gac ttc tgg tat gaa tat cga agt cca gtt caa      150
Ala Ser Ile Gly Thr Asp Phe Trp Tyr Glu Tyr Arg Ser Pro Val Gln
          5                      10                      15                      20
gaa aat tcc agt gat ttg aat aaa agc atc tgg gat gaa ttc att agt      198
Glu Asn Ser Ser Asp Leu Asn Lys Ser Ile Trp Asp Glu Phe Ile Ser
          25                      30                      35
gat gag gca gat gaa aag act tat aat gat gca ctt ttt cga tac aat      246
Asp Glu Ala Asp Glu Lys Thr Tyr Asn Asp Ala Leu Phe Arg Tyr Asn
          40                      45                      50
ggc aca gtg gga ttg tgg gga cgg tgt atc acc ata ccc aaa aac atg      294
Gly Thr Val Gly Leu Trp Gly Arg Cys Ile Thr Ile Pro Lys Asn Met
          55                      60                      65
gat tgg tat agc cca cca gaa agg aca ggt att tct ctt att tta act      342
His Trp Tyr Ser Pro Pro Glu Arg Thr Gly Ile Ser Leu Ile Leu Thr
          70                      75                      80
tct gtc ttc ttc acc tgg tta ata ata gac aaa acg acg taatgattgc      391
Ser Val Phe Phe Thr Trp Leu Ile Ile Asp Lys Thr Thr
          85                      90                      95
ccaattacat gtaagcaggt ttgttggttc tctctctcct taaagaaata aatcgtgtat      451
cttctctttc tactgccttc tctccccaac ttctttgcat taccatggta ctcatcaata      511
ttggttggat gaggaacttt tcttatcttg ggaaagcctt aatggctttt ttttttctta      571
tttactcact cattaaaata cttttcatta ctctaacaca tgttataaag aaatagttgg      631
aaaagtgcac cgaaagactt ttaaaaatat ttggtaacta gtaaaaggac taccatcgaa      691
aatcaactca aaaaattgtc cttttatggg ttagctgtat tataatacat atctatcatt      751
tgccctctgtg tcttagagga tataatttga ccagctctac atttaatctg tgtaattatg      811
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gttaaaaaaa aaaaaa
                                         886

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<210> 138  
<211> 1244  
<212> DNA



<213> Homo sapiens

<220>

<221> CDS

<222> 46..579

<221> sig\_peptide

<222> 46..156

<223> Von Heijne matrix

score 3.5

seq LVFNFLILITILT/IW

<400> 138

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cccttatcca ggtntttatc tanggaatcc cnnnaagact gggga atg gag aga cag      57
                                         Met Glu Arg Gln
                                         -35
tca agg gtt atg tca gaa aag gat gag tat cag ttt caa cat can nna      105
Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe Gln His Xaa Xaa
          -30                      -25                      -20
gcg gng gan ctg ctt gtc ttc aat ttt ttg ctc atc ctt acc att ttg      153
Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile Leu Thr Ile Leu
          -15                      -10                      -5
aca atc tgg tta ttt aaa aat cat cga ttc cgc ttc ttg cat gaa act      201
Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe Leu His Glu Thr
          1                      5                      10                      15
gga gga gca atg gtg tat ggc ctt ata atg gga cta att tca cga tat      249
Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu Ile Ser Arg Tyr
          20                      25                      30
gct aca gca cca act gat att gaa agt gga act gtc tgt gac tgt gta      297
Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val Cys Asp Cys Val
          35                      40                      45
aaa cta act ttc agt cca cca act ctg ctg gtt aat gtc act gac caa      345
Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn Val Thr Asp Gln
          50                      55                      60
gtt tat gaa tat aaa tac aaa aga gaa ata agt cag cac aac atc aat      393
Val Tyr Glu Tyr Lys Tyr Lys Arg Glu Ile Ser Gln His Asn Ile Asn
          65                      70                      75
cct cat caa gga aat gct ata ctt gaa aag atg aca ttt gat cca gaa      441
Pro His Gln Gly Asn Ala Ile Leu Glu Lys Met Thr Phe Asp Pro Glu
          80                      85                      90                      95
atc ttc ttc aat gtt tta ctg cca cca att ata ttt cat gca gga tat      489
Ile Phe Phe Asn Val Leu Leu Pro Pro Ile Ile Phe His Ala Gly Tyr
          100                      105                      110
agt cta aag aag aga cac ttt ttt caa aac tta gga tct att tta acg      537
Ser Leu Lys Lys Arg His Phe Phe Gln Asn Leu Gly Ser Ile Leu Thr
          115                      120                      125
tat gcc ttc ttg gga act gcc atc tcc tgc atc gtc ata ggg      579
Tyr Ala Phe Leu Gly Thr Ala Ile Ser Cys Ile Val Ile Gly
          130                      135                      140
taagtgcact tcggagctca agttgcaggt ggctgtgggg tctgtgatct gtgtgaggga      639
tctaacactt ccaggattct tgctggctgg gaaaattgtc ttttttttag tatatcacat      699
at ttgtatgt tttttctgac ttaattccac ggcttctgac aaatacaagg cttcaaatca      759
aagcaaaacta gaggattgct ggactttctc tgtgagttct ggacttctga cttagggaat      819
gtggatcact tgctttgagt tatgtgaagc gcattgcatt cttcttttag tttgagtaat      879
gccgatatgg tcaactgcatt cttttttgtc ttgtattgag agaccttacc tgtatttggc      939
aggagtgcaa aagtaactat atgccaagag ttttctttct aaaggaaagt ttacaagaca      999
gcagtctgaa acagatatgn tccaatatn naacagagtt gcttaataca gggatagctt      1059
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ttcagttaat accctgtaga atgcagactc tttntttcat tgtattttct tgattatgct 1119
actgagccct aagtcacacg ttatatactc tggcttgcag ctcatacataa agtaaaatgt 1179
ggtagccaaat ggtgaaggca atccagcctn tgataatccc gtccaataca ttaaagntcc 1239
actgc 1244
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<210> 139  
<211> 471  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 92..469

<221> sig\_peptide  
<222> 92..172  
<223> Von Heijne matrix  
score 7.9  
seq VVVLALGFLGCGY/AK

<221> polyA\_signal  
<222> 454..459

<221> polyA\_site  
<222> 458..471

<400> 139  
gcaagtgcag aagtcgggtga cgggtgggcat ctgggtgtca atcgatgggg catcctttct 60  
gaagatcttc gggccactgt cgtccagtgc c atg cag ttt gtc aac gtg ggc 112  
Met Gln Phe Val Asn Val Gly  
-25  
tac ttc ctc atc gca gcc ggc gtt gtg gtc ctt gct ctt ggt ttc ctg 160  
Tyr Phe Leu Ile Ala Gly Val Val Val Ala Leu Gly Phe Leu  
20 -15 -10 -5  
ggc tgc tat ggt gct aag act gag agc atg tgt gcc ctc gtg acg ttc 208  
Gly Cys Tyr Gly Ala Lys Thr Glu Ser Met Cys Ala Leu Val Thr Phe  
1 5 10  
ttc ttc atc ctc ctc ctc atc ttc att gct gag gtt gca gct gct gtg 256  
Phe Phe Ile Leu Leu Leu Ile Phe Ile Ala Glu Val Ala Ala Ala Val  
15 20 25  
gtc gcc ctg gtg tac acc aca atg gct gag cac ttc ctg acg ttg ctg 304  
Val Ala Leu Val Tyr Thr Thr Met Ala Glu His Phe Leu Thr Leu Leu  
30 35 40  
gta gtg cct gcc atc aag aaa gat tat ggt tcc cag gaa gac ttc act 352  
Val Val Pro Ala Ile Lys Lys Asp Tyr Gly Ser Gln Glu Asp Phe Thr  
45 50 55 60  
caa gtg tgg aac acc acc atg aaa ggg ctc aag tgc cgt ggc ttc acc 400  
Gln Val Trp Asn Thr Thr Met Lys Gly Leu Lys Cys Arg Gly Phe Thr  
65 70 75  
aac tat acg gat ttt gag gac tca ccc tac ttc aaa atg cat aaa cct 448  
Asn Tyr Thr Asp Phe Glu Asp Ser Pro Tyr Phe Lys Met His Lys Pro  
80 85 90  
gtt aca atg aaa aaa aaa aa 471  
Val Thr Met Lys Lys Lys Lys  
95

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<210> 140
<211> 849
<212> DNA
<213> Homo sapiens
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<220>  
<221> CDS  
<222> 154..675
```

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<221> sig_peptide
<222> 154..498
<223> Von Heijne matrix
      score 4.8
      seq PLRLLNLLILIEG/GV
```

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<221> polyA_signal
<222> 819..824
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```
<221> polyA_site
<222> 838..849
```

$$X_{400} > 140$$

cccttatctc	cagacctcat	tcgcaatgaa	gtagaatgtc	tgaagcaga	tttcaaccac	60
agaatcaagg	aggtttctct	caactccctc	ttcagtgcc	actatgttgc	attttctcccc	120
ctgtgttttg	tgaagagtac	ccagtactat	gac atg cgc	tgg tca tgt	gag cac	174
			Met Arg Trp Ser Cys Glu His			
			-115		-110	
ctc gtt atg	gtg tgg atc	aat gct ttt	gtc atg ctc	acc acg caa	ctg	222
Leu Val Met	Val Trp Ile	Asn Ala Phe	Val Met Leu	Thr Thr Gln	Leu	
	-105		-100		-95	
ttg cca tcc	aaa tac tgt	gat ttg cta	cat aaa tca	gct gct cac	ctg	270
Leu Pro Ser	Lys Tyr Cys	Asp Leu Leu	His Lys Ser	Ala Ala His	Leu	
	-90		-85		-80	
ggc aag tgg	cag aag ttg	gaa cat ggg	tcc tac agc	aat gct cca	cag	318
Gly Lys Trp	Gln Lys Leu	Glu His Gly	Ser Tyr Ser	Asn Ala Pro	Gln	
	-75		-70		-65	
cac att tgg	tca gaa aat	aca ata tgg	cct caa ggg	gtg ctg gtg	cgg	366
His Ile Trp	Ser Glu Asn	Thr Ile Trp	Pro Gln Gly	Val Leu Val	Arg	
	-60		-55		-50	
cac agc aga	tgt tta tat	aga gcc atg	ggg cct tac	aac gtg gca	gtg	414
His Ser Arg	Cys Leu Tyr	Arg Ala Met	Gly Pro Tyr	Asn Val Ala	Val	
	-40		-35		-30	
cct tca gat	gta tct cat	gcc cgc ttt	tat ttc tta	ttt cat cga	cca	462
Pro Ser Asp	Val Ser His	Ala Arg Phe	Tyr Phe Leu	Phe His Arg	Pro	
	-25		-20		-15	
tta agg ctg	tta aat ctg	ctc atc ctt	att gag ggc	ggt gtc gtc	ttc	510
Leu Arg Leu	Leu Asn Leu	Leu Ile Leu	Ile Glu Gly	Gly Val Val	Phe	
	-10		-5		1	
tat cag ctc	tat tcc ttg	ctg cgg tcg	gag aag tgg	aac cac aca	ctt	558
Tyr Gln Leu	Tyr Ser Leu	Leu Arg Ser	Glu Lys Trp	Asn His Thr	Leu	
5		10		15	20	
tcc atg gct	ctc atc ctc	ttc tgc aac	tac tat gtt	tta ttt aaa	ctt	606
Ser Met Ala	Leu Ile Leu	Phe Cys Asn	Tyr Tyr Val	Leu Phe Lys	Leu	
	25		30		35	

```
ctc cgg gac aga ata gta tta ggc agg gca tac tcc tac cca ctc aac      654
Leu Arg Asp Arg Ile Val Leu Gly Arg Ala Tyr Ser Tyr Pro Leu Asn
      40                      45                      50
agt tat gaa ctc aag gca aac taagctgcct ctcaacaatg agggagaact      705
Ser Tyr Glu Leu Lys Ala Asn
      55
cagataaaaa tattttcata cgttctatattt ttttcttgtg atttttataa atatttaaga      765
tgttttatat tttgtataact attatgtttt gaaagtcggg aagagtaagg gatattaaat      825
gtatccgtaa acaaaaaaaaaa aaaa      849
```

<210> 141  
<211> 155  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -31..-1

<400> 141

```
Met Phe Thr Ser Thr Gly Ser Ser Gly Leu Tyr Lys Ala Pro Leu Ser
-30                      -25                      -20
Lys Ser Leu Leu Leu Val Pro Ser Ala Leu Ser Leu Leu Leu Ala Leu
15                      -10                      -5                      1
Leu Leu Pro His Cys Gln Lys Pro Phe Val Tyr Asp Leu His Ala Val
5                      10                      15
Lys Asn Asp Phe Gln Ile Trp Arg Leu Ile Cys Gly Arg Ile Ile Cys
20                      25                      30
Leu Asp Leu Lys Asp Thr Phe Cys Ser Ser Leu Leu Ile Tyr Asn Phe
35                      40                      45
Arg Ile Phe Glu Arg Arg Tyr Gly Ser Arg Lys Phe Ala Ser Phe Leu
50                      55                      60                      65
Leu Gly Thr Trp Val Leu Ser Ala Leu Phe Asp Phe Leu Leu Ile Glu
70                      75                      80
Ala Met Gln Tyr Phe Phe Gly Ile Thr Ala Ala Ser Asn Leu Pro Ser
85                      90                      95
Gly Leu Ile Phe Cys Cys Ala Phe Cys Ser Glu Thr Lys Leu Phe Leu
100                      105                      110
Ser Arg Gln Ala Met Ala Glu Asn Phe Ser Ile
115                      120
```

<210> 142  
<211> 55  
<212> PRT  
<213> Homo sapiens

<400> 142

```
Met Ala Asp Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Gln Lys Arg
1                      5                      10                      15
Met Tyr Tyr Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile Thr Phe
20                      25                      30
Phe Met Gly Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu Lys Gln
35                      40                      45
```

Lys Gln Lys Lys Arg Ser Asn  
50 55

<210> 143  
<211> 67  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -20..-1

<400> 143  
Met Ser Arg Asn Leu Arg Thr Ala Leu Ile Phe Gly Gly Phe Ile Ser  
-20 -15 -10 -5  
Leu Ile Gly Ala Ala Phe Tyr Pro Ile Tyr Phe Arg Pro Leu Met Arg  
1 5 10  
Leu Glu Glu Tyr Lys Lys Glu Gln Ala Ile Asn Arg Ala Gly Ile Val  
15 20 25  
Gln Glu Asp Val Gln Pro Pro Gly Leu Lys Val Trp Ser Asp Pro Phe  
30 35 40  
Gly Arg Lys  
45

<210> 144  
<211> 198  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -21..-1

<400> 144  
Met Pro Val Pro Ala Leu Cys Leu Leu Trp Ala Leu Ala Met Val Thr  
-20 -15 -10  
Arg Pro Ala Ser Ala Ala Pro Met Gly Gly Pro Glu Leu Ala Gln His  
-5 1 5 10  
Glu Glu Leu Thr Leu Leu Phe His Gly Thr Leu Gln Leu Gly Gln Ala  
15 20 25  
Leu Asn Gly Val Tyr Arg Thr Thr Glu Gly Trp Leu Thr Lys Ala Arg  
30 35 40  
Asn Ser Leu Gly Leu Tyr Gly Arg Thr Ile Glu Leu Leu Gly Gln Glu  
45 50 55  
Val Ser Arg Gly Arg Asp Ala Ala Gln Glu Leu Arg Ala Ser Leu Leu  
60 65 70 75  
Glu Thr Gln Met Glu Glu Asp Ile Leu Gln Leu Gln Ala Glu Ala Thr  
80 85 90  
Ala Glu Val Leu Gly Glu Val Ala Gln Ala Gln Lys Val Leu Arg Asp  
95 100 105  
Ser Val Gln Arg Leu Glu Val Gln Leu Arg Ser Ala Trp Leu Gly Pro  
110 115 120  
Ala Tyr Arg Glu Phe Glu Val Leu Lys Ala His Ala Asp Lys Gln Ser

[illegible]

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<210> 145
<211> 135
<212> PRT
<213> Homo sapiens
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<220>  
<221> SIGNAL  
<222> -25..-1
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<400>	145														
Met	Ser	Leu	Arg	Asn	Leu	Trp	Arg	Asp	Tyr	Lys	Val	Leu	Val	Val	Met
-25					-20					-15					-10
Val	Pro	Leu	Val	Gly	Leu	Ile	His	Leu	Gly	Trp	Tyr	Arg	Ile	Lys	Ser
				-5					1				5		
Ser	Pro	Val	Phe	Gln	Ile	Pro	Lys	Asn	Asp	Asp	Ile	Pro	Glu	Gln	Asp
		10					15					20			
Ser	Leu	Gly	Leu	Ser	Asn	Leu	Gln	Lys	Ser	Gln	Ile	Gln	Gly	Lys	Xaa
	25					30					35				
Ala	Gly	Leu	Gln	Ser	Ser	Gly	Lys	Glu	Ala	Ala	Leu	Asn	Leu	Ser	Phe
40					45					50					55
Ile	Ser	Lys	Glu	Glu	Met	Lys	Asn	Thr	Ser	Trp	Ile	Arg	Lys	Asn	Trp
				60					65					70	
Leu	Leu	Val	Ala	Gly	Ile	Ser	Phe	Ile	Gly	Asp	His	Leu	Gly	Thr	Tyr
			75					80					85		
Phe	Leu	Gln	Arg	Ser	Ala	Lys	Gln	Ser	Val	Lys	Phe	Gln	Ser	Gln	Ser
		90					95					100			
Lys	Gln	Lys	Ser	Ile	Glu	Glu									
	105					110									

```
<210> 146
<211> 255
<212> PRT
<213> Homo sapiens
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```
<220>
<221> SIGNAL
<222> -70..-1
```

<400> 146															
Met	Gln	Gln	Lys	Glu	Gln	Gln	Phe	Arg	Glu	Trp	Phe	Leu	Lys	Glu	Phe
-70					-65					-60					-55
Pro	Gln	Ile	Arg	Trp	Lys	Ile	Gln	Glu	Ser	Ile	Glu	Arg	Leu	Arg	Val
				-50					-45						-40
Ile	Ala	Asn	Glu	Ile	Glu	Lys	Val	His	Arg	Gly	Cys	Val	Ile	Ala	Asn
			-35					-30						-25	

Val	Val	Ser	Gly	Ser	Thr	Gly	Ile	Leu	Ser	Val	Ile	Gly	Val	Met	Leu
		-20					-15					-10			
Ala	Pro	Phe	Thr	Ala	Gly	Leu	Ser	Leu	Ser	Ile	Thr	Ala	Ala	Gly	Val
	-5					1				5					10
Gly	Leu	Gly	Ile	Ala	Ser	Ala	Thr	Ala	Gly	Ile	Ala	Ser	Ser	Ile	Val
			15						20					25	
Glu	Asn	Thr	Tyr	Thr	Arg	Ser	Ala	Glu	Leu	Thr	Ala	Ser	Arg	Leu	Thr
			30					35					40		
Ala	Thr	Ser	Thr	Asp	Gln	Leu	Glu	Ala	Leu	Arg	Asp	Ile	Leu	His	Asp
		45					50					55			
Ile	Thr	Pro	Asn	Val	Leu	Ser	Phe	Ala	Leu	Asp	Phe	Asp	Glu	Ala	Thr
	60					65				70					
Lys	Met	Ile	Ala	Asn	Asp	Val	His	Thr	Leu	Arg	Arg	Ser	Lys	Ala	Thr
	75				80					85					90
Val	Gly	Arg	Pro	Leu	Ile	Ala	Trp	Arg	Tyr	Val	Pro	Ile	Asn	Val	Val
				95					100					105	
Glu	Thr	Leu	Arg	Thr	Arg	Gly	Ala	Pro	Thr	Arg	Ile	Val	Arg	Lys	Val
			110					115					120		
Ala	Arg	Asn	Leu	Gly	Lys	Ala	Thr	Ser	Gly	Val	Leu	Val	Val	Leu	Asp
		125					130				135				
Val	Val	Asn	Leu	Val	Gln	Asp	Ser	Leu	Asp	Leu	His	Lys	Gly	Glu	Lys
	140					145					150				
Ser	Glu	Ser	Ala	Glu	Leu	Leu	Arg	Gln	Trp	Ala	Gln	Glu	Leu	Glu	Glu
	155				160				165						170
Asn	Leu	Asn	Glu	Leu	Thr	His	Ile	His	Gln	Ser	Leu	Lys	Ala	Gly	
				175					180					185	

<210> 147  
 <211> 59  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -49...-1

Met	Pro	Gly	Thr	Glu	Val	Leu	Glu	Gly	Ala	Thr	Asp	Gly	Leu	Ala	Ala
				-45					-40					-35	
Ile	Asn	Leu	Leu	Lys	Trp	Ile	Lys	Thr	Leu	Gly	Gly	Ser	Val	Ile	Ser
			-30					-25					-20		
Met	Ile	Val	Leu	Leu	Ile	Cys	Val	Val	Cys	Leu	Tyr	Ile	Val	Cys	Arg
		-15				-10					-5				
Cys	Gly	Ser	His	Leu	Trp	Arg	Glu	Ser	His	His					
	1				5					10					

<210> 148  
 <211> 180  
 <212> PRT  
 <213> Homo sapiens

<400> 148  
 Met Cys Ile Ser Gly Leu Cys Gln Ile Val Gly Cys Asp His Gln Leu

1					5					10					15				
Gly	Ser	Thr	Val	Lys	Glu	Asp	Asn	Cys	Gly	Val	Cys	Asn	Gly	Asp	Gly				
			20				25						30						
Ser	Thr	Cys	Arg	Leu	Val	Arg	Gly	Gln	Tyr	Lys	Ser	Gln	Leu	Ser	Ala				
			35				40						45						
Thr	Lys	Ser	Asp	Asp	Thr	Val	Val	Ala	Ile	Pro	Tyr	Gly	Ser	Arg	His				
			50				55						60						
Ile	Arg	Leu	Val	Leu	Lys	Gly	Pro	Asp	His	Leu	Tyr	Leu	Glu	Thr	Lys				
65				70						75			80						
Thr	Leu	Gln	Gly	Thr	Lys	Gly	Glu	Asn	Ser	Leu	Ser	Ser	Thr	Gly	Thr				
			85						90			95							
Phe	Leu	Val	Asp	Asn	Ser	Ser	Val	Asp	Phe	Gln	Lys	Phe	Pro	Asp	Lys				
			100			105						110							
Glu	Ile	Leu	Arg	Met	Ala	Gly	Pro	Leu	Thr	Ala	Asp	Phe	Ile	Val	Lys				
			115			120						125							
Ile	Arg	Asn	Ser	Gly	Ser	Ala	Asp	Ser	Thr	Val	Gln	Phe	Ile	Phe	Tyr				
			130			135						140							
Gln	Pro	Ile	Ile	His	Arg	Trp	Arg	Glu	Thr	Asp	Phe	Phe	Pro	Cys	Ser				
145				150						155			160						
Ala	Thr	Cys	Gly	Gly	Tyr	Gln	Leu	Thr	Ser	Ala	Glu	Cys	Tyr	Asp					
			165						170			175							
Leu	Arg	Ser	Asn																
			180																

$\langle 210 \rangle$	149
$\langle 211 \rangle$	162
$\langle 212 \rangle$	PRT
$\langle 213 \rangle$	Homo sapiens

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<220>
<221> SIGNAL
<222> -23..-1

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[illegible]



<210> 150  
 <211> 120  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -23...-1

<400> 150  
 Met Gly Asp Lys Ile Trp Leu Pro Phe Pro Val Leu Leu Leu Ala Ala  
                   -20                  -15                  -10  
 Leu Pro Pro Val Leu Leu Pro Gly Ala Ala Gly Phe Thr Pro Ser Leu  
                   -5                  1                  5  
 Asp Ser Asp Phe Thr Phe Thr Leu Pro Ala Gly Gln Lys Glu Cys Phe  
 10                  15                  20                  25  
 Tyr Gln Pro Met Pro Leu Lys Ala Ser Leu Glu Ile Glu Tyr Gln Val  
                   30                  35                  40  
 Leu Asp Gly Ala Gly Leu Asp Ile Asp Phe His Leu Ala Ser Pro Glu  
                   45                  50                  55  
 Gly Lys Thr Leu Val Phe Glu Gln Arg Lys Ser Asp Gly Val His Thr  
                   60                  65                  70  
 Cys Ile Arg Ser Lys Asn Gly Pro Gly Thr Ala Val His Ala Tyr Asn  
                   75                  80                  85  
 Pro Ser Thr Phe Arg Gly Gln Val  
 90                  95

<210> 151  
 <211> 7  
 <212> PRT  
 <213> Homo sapiens

<400> 151  
 Met Val Glu Met Thr Gly Val  
 1                  5

<210> 152  
 <211> 199  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -42...-1

<400> 152  
 Met Asp Gly Gln Lys Lys Asn Trp Lys Asp Lys Val Val Asp Leu Leu  
                   -40                  -35                  -30  
 Tyr Trp Arg Asp Ile Lys Lys Thr Gly Val Val Phe Gly Ala Ser Leu  
                   -25                  -20                  -15

```

Phe Leu Leu Leu Ser Leu Thr Val Phe Ser Ile Val Ser Val Thr Ala
-10          -5          1          5
Tyr Ile Ala Leu Ala Leu Leu Ser Val Thr Ile Ser Phe Arg Ile Tyr
          10          15          20
Lys Gly Val Ile Gln Ala Ile Gln Lys Ser Asp Glu Gly His Pro Phe
          25          30          35
Arg Ala Tyr Leu Glu Ser Glu Val Ala Ile Ser Glu Glu Leu Val Gln
          40          45          50
Lys Tyr Ser Asn Ser Ala Leu Gly His Val Asn Cys Thr Ile Lys Glu
55          60          65          70
Leu Arg Arg Leu Phe Leu Val Asp Asp Leu Val Asp Ser Leu Lys Phe
          75          80          85
Ala Val Leu Met Trp Val Phe Thr Tyr Val Gly Ala Leu Phe Asn Gly
          90          95          100
Leu Thr Leu Leu Ile Leu Ala Leu Ile Ser Leu Phe Ser Val Pro Val
          105          110          115
Ile Tyr Glu Arg His Gln Ala Gln Ile Asp His Tyr Leu Val Leu Ala
          120          125          130
Asn Lys Asn Val Lys Asp Ala Met Ala Lys Ile Gln Ala Lys Ile Pro
135          140          145          150
Gly Leu Lys Arg Lys Ala Glu
          155

```

```

<210> 153
<211> 43
<212> PRT
<213> Homo sapiens

```

```

<400> 153
Met Pro Phe Arg Met Ser Gly Tyr Ile Pro Phe Gly Thr Pro Ile Val
          5          10          15
Ser Val Thr Phe Lys Gly Phe Pro Phe Leu Lys Asn Tyr Phe Lys Cys
          20          25          30
Leu Thr Leu Cys Tyr Cys Ser Arg Val Phe Asp
          35          40

```

```

<210> 154
<211> 50
<212> PRT
<213> Homo sapiens

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<220>
<221> SIGNAL
<222> -37..-1

```

```

<400> 154
Met Glu Trp Ala Gly Lys Gln Arg Asp Phe Gln Val Arg Ala Ala Pro
          -35          -30          -25
Gly Trp Asp His Leu Ala Ser Phe Pro Gly Pro Ser Leu Arg Leu Phe
          -20          -15          -10
Ser Gly Ser Gln Ala Ser Val Cys Ser Leu Cys Ser Gly Phe Gly Ala
-5          1          5          10
Gln Glu

```

<210> 155  
 <211> 153  
 <212> PRT  
 <213> Homo sapiens

<400> 155  
 Thr Val Pro Leu Leu Leu Glu Pro Ala Asp His Ala Arg Gly Arg Ala  
 1 5 10 15  
 His Val His Leu Pro Glu Asn Val Arg Ser Gln Ser Pro Gly His Val  
 20 25 30  
 Arg Arg Gly Arg Ser Gly Ala Gln Val Leu Pro Thr Gly Pro Asp Glu  
 35 40 45  
 Lys Gln Val Glu Lys Ser Glu Val Asp Phe Ser Lys Ser His Ser Leu  
 50 55 60  
 Val Arg Arg Phe Glu Asp Leu Lys Pro Lys Leu Ser Val Cys Lys Thr  
 65 70 75 80  
 Gly Ser Gln Val Phe Arg Ser Glu Asn Trp Lys Val Trp Ala Glu Ser  
 85 90 95  
 Ser Arg Gly Asp His Asp Asp Cys Leu Asp Leu Cys Ser Val Leu Cys  
 100 105 110  
 Trp Gly Glu Leu Leu Arg Thr Ile Pro Glu Ile Pro Pro Lys Arg Gly  
 115 120 125  
 Glu Leu Lys Thr Glu Leu Leu Gly Leu Lys Glu Arg Lys His Lys Pro  
 130 135 140  
 Gln Val Ser Gln Gln Glu Glu Leu Lys  
 145 150

<210> 156  
 <211> 67  
 <212> PRT  
 <213> Homo sapiens

<400> 156  
 Met Arg Gln Lys Arg Lys Gly Asp Leu Ser Pro Ala Lys Leu Met Met  
 1 5 10 15  
 Leu Thr Ile Gly Asp Val Ile Lys Gln Leu Ile Glu Ala His Glu Gln  
 20 25 30  
 Gly Lys Asp Ile Asp Leu Asn Lys Val Arg Thr Lys Thr Ala Ala Lys  
 35 40 45  
 Tyr Gly Leu Ser Ala Gln Pro Arg Leu Val Asp Ile Ile Ala Ala Val  
 50 55 60  
 Pro Pro Glu  
 65

<210> 157  
 <211> 87  
 <212> PRT  
 <213> Homo sapiens

<400> 157

Met Asp Glu Leu Ser Glu Glu Asp Lys Leu Thr Val Ser Arg Ala Arg  
1 5 10 15  
Lys Ile Gln Arg Phe Leu Ser Gln Pro Phe Gln Val Ala Glu Val Phe  
20 25 30  
Thr Gly His Met Gly Lys Leu Val Pro Leu Lys Glu Thr Ile Lys Gly  
35 40 45  
Phe Gln Gln Ile Leu Ala Gly Glu Tyr Asp His Leu Pro Glu Gln Ala  
50 55 60  
Phe Tyr Met Val Gly Pro Ile Glu Glu Ala Val Ala Lys Ala Asp Lys  
65 70 75 80  
Leu Ala Glu Glu His Ser Ser  
85

<210> 158

<211> 250

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -85..-1

<400> 158

Met Ser Ala Glu Val Lys Val Thr Gly Gln Asn Gln Glu Gln Phe Leu  
85 -80 -75 -70  
Leu Leu Ala Lys Ser Ala Lys Gly Ala Ala Leu Ala Thr Leu Ile His  
-65 -60 -55  
Gln Val Leu Glu Ala Pro Gly Val Tyr Val Phe Gly Glu Leu Leu Asp  
-50 -45 -40  
Met Pro Asn Val Arg Glu Leu Xaa Ala Arg Asn Leu Pro Pro Leu Thr  
-35 -30 -25  
Glu Ala Gln Lys Asn Lys Leu Arg His Leu Ser Val Val Thr Leu Ala  
-20 -15 -10  
Ala Lys Val Lys Cys Ile Pro Tyr Ala Val Leu Leu Glu Ala Leu Ala  
-5 1 5 10  
Leu Arg Asn Val Arg Gln Leu Glu Asp Leu Val Ile Glu Ala Val Tyr  
15 20 25  
Ala Asp Val Leu Arg Gly Ser Leu Asp Gln Arg Asn Gln Arg Leu Glu  
30 35 40  
Val Asp Tyr Ser Ile Gly Arg Asp Ile Gln Arg Gln Asp Leu Ser Ala  
45 50 55  
Ile Ala Arg Thr Leu Gln Glu Trp Cys Val Gly Cys Glu Val Val Leu  
60 65 70 75  
Ser Gly Ile Glu Glu Gln Val Ser Arg Ala Asn Gln His Lys Glu Gln  
80 85 90  
Gln Leu Gly Leu Lys Gln Gln Ile Glu Ser Glu Val Ala Asn Leu Lys  
95 100 105  
Lys Thr Ile Lys Val Thr Thr Ala Ala Ala Ala Thr Ser Gln  
110 115 120  
Asp Pro Glu Gln His Leu Thr Glu Leu Arg Glu Pro Ala Pro Gly Thr  
125 130 135  
Asn Gln Arg Gln Pro Ser Lys Lys Ala Ser Lys Gly Lys Gly Leu Arg  
140 145 150 155  
Gly Ser Ala Lys Ile Trp Ser Lys Ser Asn  
160 165

<210> 159  
 <211> 24  
 <212> PRT  
 <213> Homo sapiens

<400> 159  
 Met Pro Thr Asn Cys Ala Ala Ala Gly Cys Ala Thr Thr Tyr Asn Lys  
 1 5 10 15  
 His Ile Asn Ile Ser Phe His Arg  
 20

<210> 160  
 <211> 228  
 <212> PRT  
 <213> Homo sapiens

<400> 160  
 Met Pro Thr Asn Cys Ala Ala Ala Gly Cys Ala Thr Thr Tyr Asn Lys  
 1 5 10 15  
 His Ile Asn Ile Ser Phe His Arg Phe Pro Leu Asp Pro Lys Arg Arg  
 20 25 30  
 Lys Glu Trp Val Arg Leu Val Arg Arg Lys Asn Phe Val Pro Gly Lys  
 35 40 45  
 His Thr Phe Leu Cys Ser Lys His Phe Glu Ala Ser Cys Phe Asp Leu  
 50 55 60  
 Thr Gly Gln Thr Arg Arg Leu Lys Met Asp Ala Val Pro Thr Ile Phe  
 65 70 75 80  
 Asp Phe Cys Thr His Ile Lys Ser Met Lys Leu Lys Ser Arg Asn Leu  
 85 90 95  
 Leu Lys Lys Asn Asn Ser Cys Ser Pro Ala Gly Pro Ser Ser Leu Lys  
 100 105 110  
 Ser Asn Ile Ser Ser Gln Gln Val Leu Leu Glu His Ser Tyr Ala Phe  
 115 120 125  
 Arg Asn Pro Met Glu Ala Lys Lys Arg Ile Ile Lys Leu Glu Lys Glu  
 130 135 140  
 Ile Ala Ser Leu Arg Arg Lys Met Lys Thr Cys Leu Gln Lys Glu Arg  
 145 150 155 160  
 Arg Ala Thr Arg Arg Trp Ile Lys Ala Met Cys Leu Val Lys Asn Leu  
 165 170 175  
 Glu Ala Asn Ser Val Leu Pro Lys Gly Thr Ser Glu His Met Leu Pro  
 180 185 190  
 Thr Ala Leu Ser Ser Leu Pro Leu Glu Asp Phe Lys Ile Leu Glu Gln  
 195 200 205  
 Asp Gln Gln Asp Lys Thr Leu Leu Ser Leu Asn Leu Lys Gln Thr Lys  
 210 215 220  
 Ser Thr Phe Ile  
 225

<210> 161  
 <211> 86

<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -20..-1

<400> 161  
Met Asn Leu His Phe Pro Gln Trp Phe Val His Ser Ser Ala Leu Gly  
-20 -15 -10 -5  
Leu Val Leu Ala Pro Pro Phe Ser Ser Pro Gly Thr Asp Pro Thr Phe  
1 5 10  
Pro Cys Ile Tyr Cys Arg Leu Leu Asn Met Ile Met Thr Arg Leu Ala  
15 20 25  
Phe Ser Phe Ile Thr Cys Leu Cys Pro Asn Leu Lys Glu Val Cys Leu  
30 35 40  
Ile Leu Pro Glu Lys Asn Cys Asn Ser Arg His Ala Gly Phe Val Gly  
45 50 55 60  
Pro Ala Lys Leu Arg Gln  
65

<210> 162  
<211> 44  
<212> PRT  
<213> Homo sapiens  
<400> 162  
Met Ser Pro Arg Leu Glu Cys Ser Gly Ala Ile Leu Ala His Cys Asn  
5 10 15  
Pro Arg Leu Pro Gly Ser Ser Tyr Ser Pro Ala Ser Ala Thr Trp Val  
20 25 30  
Arg Gly Ser Leu Glu Pro Gly Arg Leu Arg Leu Gln  
35 40

<210> 163  
<211> 314  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -58..-1

<400> 163  
Met Gln Asn Val Ile Asn Thr Val Lys Gly Lys Ala Leu Glu Val Ala  
-55 -50 -45  
Glu Tyr Leu Thr Pro Val Leu Lys Glu Ser Lys Phe Arg Glu Thr Gly  
-40 -35 -30  
Val Ile Thr Pro Glu Glu Phe Val Ala Ala Gly Asp His Leu Val His  
-25 -20 -15  
His Cys Pro Thr Trp Gln Trp Ala Thr Gly Glu Glu Leu Lys Val Lys  
-10 -5 1 5  
Ala Tyr Leu Pro Thr Gly Lys Gln Phe Leu Val Thr Lys Asn Val Pro

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A210> 164
A211> 89
A212> PRT
A213> Homo sapiens

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<400> 164

[illegible]

<210> 165  
 <211> 98  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -15..-1

<400> 165  
 Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu Ala Val Leu Ala Trp  
 -15 -10 -5 1  
 Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg Met Lys Ser Arg Glu  
 5 10 15  
 Gln Gly Gly Arg Leu Gly Ala Glu Ser Arg Thr Leu Leu Val Ile Ala  
 20 25 30  
 His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro Thr Val Leu Gly Leu  
 35 40 45  
 Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys Phe Ser Ala Val Phe  
 50 55 60 65  
 Arg Arg Glu Leu Ser Glu Tyr Thr Glu Gly Leu Thr Ser Glu Pro Leu  
 70 75 80  
 Thr Ala

<210> 166  
 <211> 92  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -36..-1

<400> 166  
 Met Leu Val Thr Gln Gly Leu Val Tyr Gln Gly Tyr Leu Ala Ala Asn  
 -35 -30 -25  
 Ser Arg Phe Gly Ser Leu Pro Lys Val Ala Leu Ala Gly Leu Leu Gly  
 -20 -15 -10 -5  
 Phe Gly Leu Gly Lys Val Ser Tyr Ile Gly Val Cys Gln Ser Lys Phe  
 1 5 10  
 His Phe Phe Glu Asp Gln Leu Arg Gly Ala Gly Phe Gly Pro Gln His  
 15 20 25  
 Asn Arg His Cys Leu Leu Thr Cys Glu Glu Cys Lys Ile Lys His Gly  
 30 35 40  
 Leu Ser Glu Lys Gly Asp Ser Gln Pro Ser Ala Ser  
 45 50 55

<210> 167  
 <211> 351  
 <212> PRT  
 <213> Homo sapiens



<220>

<221> SIGNAL

<222> -16..-1

<400> 167

```

Met Val Pro Phe Ile Tyr Leu Gln Ala His Phe Thr Leu Cys Ser Gly
  -15                -10                -5
Trp Ser Ser Thr Tyr Arg Asp Leu Arg Lys Gly Val Tyr Val Pro Tyr
 1                    5                10                15
Thr Gln Gly Lys Trp Glu Gly Glu Leu Gly Thr Asp Leu Val Ser Ile
                20                25                30
Pro His Gly Pro Asn Val Thr Val Arg Ala Asn Ile Ala Ala Ile Thr
 35                40                45
Glu Ser Asp Lys Phe Phe Ile Asn Gly Ser Asn Trp Glu Gly Ile Leu
 50                55                60
Gly Leu Ala Tyr Ala Glu Ile Ala Arg Pro Asp Asp Ser Pro Glu Pro
 65                70                75                80
Phe Phe Asp Ser Leu Val Lys Gln Thr His Val Pro Asn Leu Phe Ser
                85                90                95
Leu Gln Leu Cys Gly Ala Gly Phe Pro Leu Asn Gln Ser Glu Val Leu
                100                105                110
Ala Ser Val Gly Gly Ser Met Ile Ile Gly Gly Ile Asp His Ser Leu
                115                120                125
Tyr Thr Gly Ser Leu Trp Tyr Thr Pro Ile Arg Arg Glu Trp Tyr Tyr
 130                135                140
Glu Val Ile Ile Val Arg Val Glu Ile Asn Gly Gln Asp Leu Lys Met
 145                150                155                160
Asp Cys Lys Glu Tyr Asn Tyr Asp Lys Ser Ile Val Asp Ser Gly Thr
                165                170                175
Thr Asn Leu Arg Leu Pro Lys Lys Val Phe Glu Ala Ala Val Lys Ser
                180                185                190
Ile Lys Ala Ala Ser Ser Thr Glu Lys Phe Pro Asp Gly Phe Trp Leu
                195                200                205
Gly Glu Gln Leu Val Cys Trp Gln Ala Gly Thr Thr Pro Trp Asn Ile
 210                215                220
Phe Pro Val Ile Ser Leu Tyr Leu Met Gly Glu Val Thr Asn Gln Ser
 225                230                235                240
Phe Arg Ile Thr Ile Leu Pro Gln Gln Tyr Leu Arg Pro Val Glu Asp
                245                250                255
Val Ala Thr Ser Gln Asp Asp Cys Tyr Lys Phe Ala Ile Ser Gln Ser
                260                265                270
Ser Thr Gly Thr Val Met Gly Ala Val Ile Met Glu Gly Phe Tyr Val
 275                280                285
Val Phe Asp Arg Ala Arg Lys Arg Ile Gly Phe Ala Val Ser Ala Cys
 290                295                300
His Val His Asp Glu Phe Arg Thr Ala Ala Val Glu Gly Pro Phe Cys
 305                310                315                320
His Leu Gly His Gly Arg Leu Trp Leu Gln His Ser Thr Asp Arg
                325                330                335

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<210> 168

<211> 138

<212> PRT

<213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -47..-1

<400> 168  
 Met Glu Lys Phe Val Asp Pro Gly Asn His Asn Ser Gly Ile Asp Leu  
           -45                    -40                    -35  
 Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu Leu Pro Phe Val Ser  
       -30                    -25                    -20  
 Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly Leu Cys Ala Cys Ile  
       -15                    -10                    -5                    1  
 Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly Ile Leu His Leu Leu  
           5                    10                    15  
 Ala Gly Leu Cys Thr Leu Gly Ser Val Ser Cys Tyr Val Ala Gly Ile  
           20                    25                    30  
 Glu Leu Leu His Gln Lys Leu Glu Leu Pro Asp Asn Val Ser Gly Glu  
       35                    40                    45  
 Phe Gly Trp Ser Phe Cys Leu Ala Cys Val Ser Ala Pro Leu Gln Phe  
       50                    55                    60                    65  
 Met Ala Ser Ala Leu Phe Ile Trp Ala Ala His Thr Asn Arg Arg Glu  
                     70                    75                    80  
 Tyr Thr Leu Met Lys Ala Tyr Arg Val Ala  
           85                    90

<210> 169  
 <211> 101  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -73..-1

<400> 169  
 Met Asn Leu Glu Arg Val Ser Asn Glu Glu Lys Leu Asn Leu Cys Arg  
           -70                    -65                    -60  
 Lys Tyr Tyr Leu Gly Gly Phe Ala Phe Leu Pro Phe Leu Trp Leu Val  
       -55                    -50                    -45  
 Asn Ile Phe Trp Phe Tyr Arg Glu Ala Phe Leu Val Pro Ala Tyr Thr  
       -40                    -35                    -30  
 Glu Gln Ser Gln Ile Lys Gly Tyr Val Trp Arg Ser Ala Val Gly Phe  
       -25                    -20                    -15                    -10  
 Leu Phe Trp Val Ile Val Leu Thr Ser Trp Ile Thr Ile Phe Gln Ile  
           -5                    1                    5  
 Tyr Arg Pro Arg Trp Gly Ala Leu Gly Asp Tyr Leu Ser Phe Thr Ile  
           10                    15                    20  
 Pro Leu Gly Thr Pro  
       25

<210> 170  
 <211> 252  
 <212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -68...-1

<400> 170

```

Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu
      -65                      -60                      -55
Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val
      -50                      -45                      -40
Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser
      -35                      -30                      -25
Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro
      -20                      -15                      -10                      -5
Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly
      1                      5                      10
Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His
      15                      20                      25
Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu
      30                      35                      40
Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys
      45                      50                      55                      60
Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe
      65                      70                      75
Arg Glu Asn Val Leu Arg Asn Leu Ala Asp Lys Ala Phe Asp Arg Pro
      80                      85                      90
Ile Cys Glu Ala Leu Leu Asp Gln Arg Phe Phe Asn Gly Ile Gly Asn
      95                      100                      105
Tyr Leu Arg Ala Glu Ile Leu Tyr Arg Leu Lys Ile Pro Pro Phe Glu
      110                      115                      120
Lys Ala Arg Ser Val Leu Glu Ala Leu Gln Gln His Arg Pro Ser Pro
      125                      130                      135                      140
Glu Leu Thr Leu Ser Gln Lys Ile Arg Thr Lys Leu Gln Asn Ser Asp
      145                      150                      155
Leu Leu Glu Leu Cys His Ser Val Pro Lys Glu Val Val Gln Leu Gly
      160                      165                      170
Glu Ala Lys Asp Gly Ser Asn Leu Cys Phe Ser Lys
      175                      180

```

<210> 171

<211> 350

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -68...-1

<400> 171

```

Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu
      -65                      -60                      -55
Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val
      -50                      -45                      -40
Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser

```

-35		-30		-25
Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro				
-20		-15		-10
Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly				
	1		5	10
Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His				
	15		20	25
Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu				
	30		35	40
Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys				
45		50		55
Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe				
	65		70	75
Arg Leu Lys Ile Pro Pro Phe Glu Lys Ala Arg Ser Val Leu Glu Ala				
	80		85	90
Leu Gln Gln His Arg Pro Ser Pro Glu Leu Thr Leu Ser Gln Lys Ile				
	95		100	105
Arg Thr Lys Leu Gln Asn Pro Asp Leu Leu Glu Leu Cys His Ser Val				
	110		115	120
Pro Lys Glu Val Asp Gln Leu Gly Gly Arg Gly Tyr Gly Ser Glu Ser				
125		130		135
Gly Glu Glu Asp Phe Ala Ala Phe Arg Ala Trp Leu Arg Cys Tyr Gly				
	145		150	155
Met Pro Gly Met Ser Ser Leu Gln Asp Arg His Gly Arg Thr Ile Trp				
	160		165	170
Phe Gln Gly Asp Pro Gly Pro Leu Ala Pro Lys Gly Arg Lys Ser Arg				
	175		180	185
Lys Lys Lys Ser Lys Ala Thr Gln Leu Ser Pro Glu Asp Arg Val Glu				
	190		195	200
Asp Ala Leu Pro Pro Ser Lys Ala Pro Ser Lys Thr Arg Arg Ala Lys				
205		210		215
Arg Asp Leu Pro Lys Arg Thr Ala Thr Gln Arg Pro Glu Gly Thr Ser				
	225		230	235
Leu Gln Gln Asp Pro Glu Ala Pro Thr Val Pro Lys Lys Gly Arg Arg				
	240		245	250
Lys Gly Arg Gln Ala Ala Ser Gly His Cys Arg Pro Arg Lys Val Lys				
	255		260	265
Ala Asp Ile Pro Ser Leu Glu Pro Glu Gly Thr Ser Ala Ser				
270		275		280

<210> 172

<211> 390

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -68...-1

<400> 172

Met Pro Glu Gly Pro Glu Leu His Leu Ala Ser Gln Phe Val Asn Glu			
	-65		-55
Ala Cys Arg Ala Leu Val Phe Gly Gly Cys Val Glu Lys Ser Ser Val			
	-50		-40
Ser Arg Asn Pro Glu Val Pro Phe Glu Ser Ser Ala Tyr Arg Ile Ser			

```

-35          -30          -25
Ala Ser Ala Arg Gly Lys Glu Leu Arg Leu Ile Leu Ser Pro Leu Pro
-20          -15          -10          -5
Gly Ala Gln Pro Gln Gln Glu Pro Leu Ala Leu Val Phe Arg Phe Gly
      1          5          10
Met Ser Gly Ser Phe Gln Leu Val Pro Arg Glu Glu Leu Pro Arg His
      15          20          25
Ala His Leu Arg Phe Tyr Thr Ala Pro Pro Gly Pro Arg Leu Ala Leu
      30          35          40
Cys Phe Val Asp Ile Arg Arg Phe Gly Arg Trp Asp Leu Gly Gly Lys
45          50          55          60
Trp Gln Pro Gly Arg Gly Pro Cys Val Leu Gln Glu Tyr Gln Gln Phe
      65          70          75
Arg Glu Asn Val Leu Arg Asn Leu Ala Asp Lys Ala Phe Asp Arg Pro
      80          85          90
Ile Cys Glu Ala Leu Leu Asp Gln Arg Phe Phe Asn Gly Ile Gly Asn
      95          100          105
Tyr Leu Arg Ala Glu Ile Leu Tyr Arg Leu Lys Ile Pro Pro Phe Glu
      110          115          120
Lys Ala Arg Ser Val Leu Glu Ala Leu Gln Gln His Arg Pro Ser Pro
125          130          135          140
Glu Leu Thr Leu Ser Gln Lys Ile Arg Thr Lys Leu Gln Asn Pro Asp
      145          150          155
Leu Leu Glu Leu Cys His Ser Val Pro Lys Glu Val Val Gln Leu Gly
      160          165          170
Gly Arg Gly Tyr Gly Ser Glu Ser Gly Glu Glu Asp Phe Ala Ala Phe
      175          180          185
Arg Ala Trp Leu Arg Cys Tyr Gly Met Pro Gly Met Ser Ser Leu Gln
      190          195          200
Asp Arg His Gly Arg Thr Ile Trp Phe Gln Gly Asp Pro Gly Pro Leu
205          210          215          220
Ala Pro Lys Gly Arg Lys Ser Arg Lys Lys Lys Ser Lys Ala Thr Gln
      225          230          235
Leu Ser Pro Glu Asp Arg Val Glu Asp Ala Leu Pro Pro Ser Lys Ala
      240          245          250
Pro Ser Arg Thr Arg Arg Ala Lys Arg Asp Leu Pro Lys Arg Thr Ala
      255          260          265
Thr Gln Arg Pro Glu Gly Thr Ser Leu Gln Gln Asp Pro Glu Ala Pro
      270          275          280
Thr Val Pro Lys Lys Gly Arg Arg Lys Gly Arg Gln Ala Ala Ser Gly
285          290          295          300
His Cys Arg Pro Arg Lys Val Lys Ala Asp Ile Pro Ser Leu Glu Pro
      305          310          315
Glu Gly Thr Ser Ala Ser
      320

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<210> 173

<211> 190

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -82..-1

<400> 173

```

Met Tyr Val Trp Pro Cys Ala Val Val Leu Ala Gln Tyr Leu Trp Phe
   -80                               -75                               -70
His Arg Arg Ser Leu Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala Gly
   -65                               -60                               -55
Val Ser Leu Pro Gly Ile Leu Thr Ala Lys Cys Gly Ala Glu Val Ile
   -50                               -45                               -40                               -35
Leu Ser Asp Ser Ser Glu Leu Pro His Cys Leu Glu Val Cys Arg Gln
                               -30                               -25                               -20
Ser Cys Gln Met Asn Asn Leu Pro His Leu Gln Val Val Gly Leu Thr
                               -15                               -10                               -5
Trp Gly His Ile Ser Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp Ile
   1                               5                               10
Ile Leu Ala Ser Asp Val Phe Phe Glu Pro Glu Asp Phe Glu Asp Ile
  15                               20                               25                               30
Leu Ala Thr Ile Tyr Phe Leu Met His Lys Asn Pro Lys Val Gln Leu
                               35                               40                               45
Trp Ser Thr Tyr Gln Val Arg Ser Ala Asp Trp Ser Leu Glu Ala Leu
                               50                               55                               60
Leu Tyr Lys Trp Asp Met Lys Cys Val His Ile Pro Leu Glu Ser Phe
   65                               70                               75
Asp Ala Asp Lys Glu Asp Ile Ala Glu Ser Thr Leu Pro Gly Arg His
   80                               85                               90
Thr Val Glu Met Leu Val Ile Ser Phe Ala Lys Asp Ser Leu
  95                               100                               105

```

<210> 174

<211> 285

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -232...-1

<400> 174

```

Met Gly Cys Val Phe Gln Ser Thr Glu Asp Lys Arg Ile Phe Lys Ile
   -230                               -225                               -220
Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu
   -215                               -210                               -205
Tyr Tyr Tyr Ser Asn Leu Ser Val Pro Ile Gly Arg Phe Gln Asn Arg
   -200                               -195                               -190                               -185
Val His Leu Met Gly Asp Asn Leu Cys Asn Asp Gly Ser Leu Leu Leu
                               -180                               -175                               -170
Gln Asp Val Gln Glu Ala Asp Gln Gly Thr Tyr Ile Cys Glu Ile Arg
   -165                               -160                               -155
Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val
   -150                               -145                               -140
Leu Pro Glu Glu Pro Lys Glu Leu Met Val His Val Gly Gly Leu Ile
   -135                               -130                               -125
Gln Met Gly Cys Val Phe Gln Ser Thr Glu Val Lys His Val Thr Lys
   -120                               -115                               -110                               -105
Val Glu Trp Ile Phe Ser Gly Arg Arg Ala Lys Glu Glu Ile Val Phe
                               -100                               -95                               -90
Arg Tyr Tyr His Lys Leu Arg Met Ser Ala Glu Tyr Ser Gln Ser Trp

```

				-85				-80					-75			
Gly	His	Phe	Gln	Asn	Arg	Val	Asn	Leu	Val	Gly	Asp	Ile	Phe	Arg	Asn	
		-70						-65					-60			
Asp	Gly	Ser	Ile	Met	Leu	Gln	Gly	Val	Arg	Glu	Ser	Asp	Gly	Gly	Asn	
		-55					-50					-45				
Tyr	Thr	Cys	Ser	Ile	His	Leu	Gly	Asn	Leu	Val	Phe	Lys	Lys	Thr	Ile	
					-35						-30				-25	
Val	Leu	His	Val	Ser	Pro	Glu	Glu	Pro	Arg	Thr	Leu	Val	Thr	Pro	Ala	
				-20					-15						-10	
Ala	Leu	Arg	Pro	Leu	Val	Leu	Gly	Gly	Asn	Gln	Leu	Val	Ile	Ile	Val	
			-5					1				5				
Gly	Ile	Val	Cys	Ala	Thr	Ile	Leu	Leu	Leu	Pro	Val	Leu	Ile	Leu	Ile	
		10				15					20					
Val	Lys	Lys	Thr	Cys	Gly	Asn	Lys	Ser	Ser	Val	Asn	Ser	Thr	Val	Leu	
25					30					35					40	
Val	Lys	Asn	Thr	Lys	Lys	Thr	Asn	Pro	Lys	Lys	Lys	Lys				
				45					50							

<210> 175

<211> 153

<212> PRT

<213> Homo sapiens

$\langle 400 \rangle$  175

[illegible]

<210> 176

<211> 49

<212> PRT

<213> Homo sapiens

<400> 176

Met Leu Xaa Gly Asp His Arg Ala Leu Leu Leu Lys Ile Trp Leu Leu  
1 5 10 15

Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro Gly Arg Leu Val Val  
20 25 30  
Met Glu Arg Arg Val Lys Met Thr Ser Cys Pro Ser Cys Pro Arg Phe  
35 40 45  
Cys

<210> 177  
<211> 99  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -24...-1

<400> 177  
Met Lys Ser Ala Lys Leu Gly Phe Leu Leu Arg Phe Phe Ile Phe Cys  
-20 -15 -10  
Ser Leu Asn Thr Leu Leu Leu Gly Gly Val Asn Lys Ile Ala Glu Lys  
-5 1 5  
Glu Cys Gly Asp Leu Lys Asp Pro Cys Lys Leu Asp Met Asn Phe Gly  
10 15 20  
Ser Cys Tyr Glu Val His Phe Arg Tyr Phe Tyr Asn Arg Thr Ser Lys  
25 30 35 40  
Arg Cys Glu Thr Phe Val Phe Ser Gly Cys Asn Gly Asn Leu Asn Asn  
45 50 55  
Phe Lys Leu Lys Ile Glu Arg Glu Val Ala Cys Val Ala Lys Tyr Lys  
60 65 70  
Pro Pro Arg  
75

<210> 178  
<211> 95  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -37...-1

<400> 178  
Met Ala Ser Pro Ala Val Asn Arg Trp Lys Arg Pro Arg Leu Lys Pro  
-35 -30 -25  
Val Trp Pro Arg Arg Leu Glu Ser Trp Leu Leu Leu Asp Ala Leu Leu  
-20 -15 -10  
Arg Leu Gly Asp Thr Lys Lys Lys Arg Gln Pro Glu Ala Ala Thr Lys  
-5 1 5 10  
Ser Cys Val Arg Ser Ser Cys Gly Gly Pro Ser Gly Asp Gly Pro Pro  
15 20 25  
Pro Cys Leu Gln Gln Pro Asp Pro Arg Ala Leu Ser Gln Ala Phe Ser  
30 35 40  
Arg Ser Phe Pro Leu Phe Pro Ser Leu Ala Gly Lys Ser Met Ile  
45 50 55



<210> 179  
 <211> 121  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -23..-1

<400> 179  
 Met Met Leu Pro Gln Trp Leu Leu Leu Leu Phe Leu Leu Phe Phe Phe  
                   -20                                  -15                  -10  
 Leu Phe Leu Leu Thr Arg Gly Ser Leu Ser Pro Thr Lys Tyr Asn Leu  
                   -5                                  1                                  5  
 Leu Glu Leu Lys Glu Ser Cys Ile Arg Asn Gln Asp Cys Glu Thr Gly  
 10                                  15                                  20                                  25  
 Cys Cys Gln Arg Ala Pro Asp Asn Cys Glu Ser His Cys Ala Glu Lys  
                                   30                                  35                                  40  
 Gly Ser Glu Gly Ser Leu Cys Gln Thr Gln Val Phe Phe Gly Gln Tyr  
                                   45                                  50                                  55  
 Arg Ala Cys Pro Cys Leu Arg Asn Leu Thr Cys Ile Tyr Ser Lys Asn  
                                   60                                  65                                  70  
 Glu Lys Trp Leu Ser Ile Ala Tyr Gly Arg Cys Gln Lys Ile Gly Arg  
                                   75                                  80                                  85  
 Gln Lys Leu Ala Lys Lys Met Phe Phe  
 90                                  95

<210> 180  
 <211> 59  
 <212> PRT  
 <213> Homo sapiens

<400> 180  
 Met Ile Leu Cys Phe Leu Leu Pro His His Arg Leu Gln Glu Ala Arg  
 1                                  5                                  10                                  15  
 Gln Ile Gln Val Leu Lys Met Leu Pro Arg Glu Lys Leu Arg Arg Arg  
                                   20                                  25                                  30  
 Glu Glu Arg Lys Gln Ile Asn Gly Lys Lys Glu Arg Thr Lys Tyr Glu  
                                   35                                  40                                  45  
 Thr Pro Arg Lys Arg Glu Gly Lys Lys Lys Lys  
 50                                  55

<210> 181  
 <211> 86  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -14..-1

<400> 181

```
Met Val Ala Leu Asn Leu Ile Leu Val Pro Cys Cys Ala Ala Trp Cys
          -10          -5          1
Asp Pro Arg Arg Ile His Ser Gln Asp Asp Val Pro Arg Ser Ser Ala
      5          10          15
Ala Asp Thr Gly Ser Ala Met Gln Arg Arg Glu Ala Trp Ala Gly Trp
      20          25          30
Arg Arg Ser Gln Pro Phe Ser Val Gly Leu Pro Ser Ala Glu Arg Leu
35          40          45          50
Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg Ser Leu Val Gly Glu Gly
      55          60          65
Tyr Arg Ile Cys Asp Leu
      70
```

<210> 182

<211> 165

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -58...-1

<400> 182

```
Met Thr Arg Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro Ile
          -55          -50          -45
Pro Val Pro Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser Pro
      -40          -35          -30
Val Arg Pro Pro Val Ser Thr Trp Gly Pro Ser Trp Ala Gln Leu Leu
      -25          -20          -15
Asp Ser Val Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln Ala Val
10          -5          1          5
Phe Ser Thr Thr Gly Pro Ala Leu Leu Leu Leu Val Ser Phe Leu
      10          15          20
Thr Phe Asp Leu Leu His Arg Pro Ala Gly His Thr Leu Pro Gln Arg
      25          30          35
Lys Leu Leu Thr Arg Gly Gln Ser Gln Gly Ala Gly Glu Gly Pro Gly
      40          45          50
Gln Gln Glu Ala Leu Leu Leu Gln Met Gly Thr Val Ser Gly Gln Leu
55          60          65          70
Ser Leu Gln Asp Ala Leu Leu Leu Leu Leu Met Gly Leu Gly Pro Leu
      75          80          85
Leu Arg Ala Cys Gly Met Pro Leu Thr Leu Leu Gly Leu Ala Phe Cys
      90          95          100
Leu His Pro Trp Ala
      105
```

<210> 183

<211> 80

<212> PRT

<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -35..-1

<400> 183

Met	Pro	Phe	Gln	Phe	Gly	Thr	Gln	Pro	Arg	Arg	Phe	Pro	Val	Glu	Gly
-35					-30				-25					-20	
Gly	Asp	Ser	Ser	Ile	Glu	Leu	Glu	Pro	Gly	Leu	Ser	Ser	Ser	Ala	Ala
				-15				-10						-5	
Cys	Asn	Gly	Lys	Glu	Met	Ser	Pro	Thr	Arg	Gln	Leu	Arg	Arg	Cys	Pro
		1				5					10				
Gly	Ser	His	Cys	Leu	Thr	Ile	Thr	Asp	Val	Pro	Val	Thr	Val	Tyr	Ala
15					20				25						
Thr	Thr	Arg	Lys	Pro	Pro	Ala	Gln	Ser	Ser	Lys	Glu	Met	His	Pro	Lys
30					35				40						45

<210> 184  
<211> 73  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -21..-1

<400> 184

Met	Ala	Pro	Gln	Thr	Leu	Leu	Pro	Val	Leu	Val	Leu	Cys	Val	Leu	Leu
-20					-15				-10						
Leu	Gln	Ala	Gln	Gly	Gly	Tyr	Arg	Asp	Lys	Met	Arg	Met	Gln	Arg	Ile
-5				1				5					10		
Lys	Val	Cys	Glu	Lys	Arg	Pro	Ser	Ile	Asp	Leu	Cys	Ile	His	His	Cys
			15				20					25			
Ser	Cys	Phe	Gln	Lys	Cys	Glu	Thr	Asn	Lys	Ile	Cys	Cys	Ser	Ala	Phe
		30				35					40				
Cys	Gly	Asn	Ile	Cys	Met	Ser	Ile	Leu							
45						50									

<210> 185  
<211> 98  
<212> PRT  
<213> Homo sapiens

<400> 185

Met	Leu	Gly	Ala	Glu	Thr	Glu	Glu	Lys	Leu	Phe	Asp	Ala	Pro	Leu	Ser
1			5					10						15	
Ile	Ser	Lys	Arg	Glu	Gln	Leu	Glu	Gln	Val	Pro	Glu	Asn	Tyr	Phe	
			20				25					30			
Tyr	Val	Pro	Asp	Leu	Gly	Gln	Val	Pro	Glu	Ile	Asp	Val	Pro	Ser	Tyr
	35					40					45				
Leu	Pro	Asp	Leu	Pro	Gly	Ile	Ala	Asn	Asp	Leu	Met	Tyr	Ile	Ala	Asp
50					55				60						
Leu	Gly	Pro	Gly	Ile	Ala	Pro	Ser	Ala	Pro	Gly	Thr	Ile	Pro	Glu	Leu
65					70				75						80

Pro Thr Phe His Thr Glu Val Ala Glu Pro Leu Lys Thr Tyr Lys Met  
85 90 95  
Gly Tyr

<210> 186  
<211> 112  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -21..-1

<400> 186  
Met Glu Ser Arg Val Leu Leu Arg Thr Phe Cys Leu Ile Phe Gly Leu  
-20 -15 -10  
Gly Ala Val Trp Gly Leu Gly Val Asp Pro Ser Leu Gln Ile Asp Val  
-5 1 5 10  
Leu Thr Glu Leu Glu Leu Gly Glu Ser Thr Thr Gly Val Arg Gln Val  
15 20 25  
Pro Gly Leu His Asn Gly Thr Lys Ala Phe Leu Phe Gln Asp Thr Pro  
30 35 40  
Arg Ser Ile Lys Ala Ser Thr Ala Thr Ala Glu Gln Phe Phe Gln Lys  
45 50 55  
Leu Arg Asn Lys His Glu Phe Thr Ile Leu Val Thr Leu Lys Gln Thr  
60 65 70 75  
His Leu Asn Ser Gly Val Ile Leu Ser Ile His His Leu Asp His Arg  
80 85 90

<210> 187  
<211> 70  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -44..-1

<400> 187  
Met Cys Cys Tyr Cys Arg Ile Phe Cys Leu Arg Cys Thr Tyr Phe Pro  
-40 -35 -30  
Val His Cys Gly Met Cys Asn Leu Arg Tyr Phe Glu Phe Ser Thr Phe  
-25 -20 -15  
Leu Leu Ser Leu Ser Leu Ile Thr Tyr Cys Phe Trp Asp Pro Pro His  
-10 -5 1  
Arg Gly Ser His Ser Leu Ser Leu Glu His Thr Pro Leu Asp Phe Leu  
5 10 15 20  
Glu Trp Gly Leu Leu Arg  
25

<210> 188

<211> 92  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -13...-1

<400> 188  
Met Leu Phe Ser Leu Ser Leu Leu Ser Asn Leu Asn Gln Ile Gly Ser  
                  -10                  -5                  1  
Ser His Leu Asp Arg Pro His Ile Pro Gly Gln Ser Ala Gln Leu Phe  
  5                  10                  15  
Ile Tyr Gln Met Ser Ser Gln Gln Leu Gln Gln Pro Ser Ala Asn  
20                  25                  30                  35  
Lys Lys Ala Gly Lys Ile His Asn Thr Pro Phe Ala Asn Gln Leu Asn  
                  40                  45                  50  
Pro Thr Gln His Leu Ala Lys Pro Phe Gln Gln Ile Leu Pro Gly Arg  
                  55                  60                  65  
Gln Ser Gly Ser Leu Thr Ser Pro Phe Leu Ala Cys  
      70                  75

<210> 189  
<211> 207  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -42...-1

<400> 189  
Met His Ile Leu Gln Leu Leu Thr Thr Val Asp Asp Gly Ile Gln Ala  
          -40                  -35                  -30  
Ile Val His Cys Pro Asp Thr Gly Lys Asp Ile Trp Asn Leu Leu Phe  
          -25                  -20                  -15  
Asp Leu Val Cys His Glu Phe Cys Gln Ser Asp Asp Pro Pro Ile Ile  
-10                  -5                  1                  5  
Leu Gln Glu Gln Lys Thr Val Leu Ala Ser Val Phe Ser Val Leu Ser  
          10                  15                  20  
Ala Ile Tyr Ala Ser Gln Thr Glu Gln Glu Tyr Leu Lys Ile Glu Lys  
          25                  30                  35  
Val Asp Leu Pro Leu Ile Asp Ser Leu Ile Arg Val Leu Gln Asn Met  
40                  45                  50  
Glu Gln Cys Gln Lys Lys Pro Glu Asn Ser Ala Glu Ser Asn Thr Glu  
55                  60                  65                  70  
Glu Thr Lys Arg Thr Asp Leu Thr Gln Asp Asp Leu His Leu Lys Ile  
          75                  80                  85  
Leu Lys Asp Ile Leu Cys Glu Phe Leu Ser Asn Ile Phe Gln Ala Leu  
          90                  95                  100  
Thr Lys Glu Thr Val Ala Gln Gly Val Lys Glu Gly Gln Leu Ser Lys  
105                  110                  115  
Gln Lys Cys Ser Ser Ala Phe Gln Asn Leu Leu Pro Phe Tyr Ser Pro  
120                  125                  130  
Val Val Glu Asp Phe Ile Lys Ile Leu Arg Glu Val Asp Lys Ala Leu

135		140		145		150								
Ala	Asp	Asp	Leu	Glu	Lys	Asn	Phe	Pro	Ser	Leu	Lys	Val	Gln	Thr
		155							160					165

<210> 190  
 <211> 201  
 <212> PRT  
 <213> Homo sapiens

<400> 190

Met	Gln	Val	Ala	Leu	Lys	Glu	Asp	Leu	Asp	Ala	Leu	Lys	Glu	Lys	Phe
1			5						10					15	
Arg	Thr	Met	Glu	Ser	Asn	Gln	Lys	Ser	Ser	Phe	Gln	Glu	Ile	Pro	Lys
			20					25					30		
Leu	Asn	Glu	Glu	Leu	Leu	Ser	Lys	Gln	Lys	Gln	Leu	Glu	Lys	Ile	Glu
		35					40					45			
Ser	Gly	Glu	Met	Gly	Leu	Asn	Lys	Val	Trp	Ile	Asn	Ile	Thr	Glu	Met
	50					55					60				
Asn	Lys	Gln	Ile	Ser	Leu	Leu	Thr	Ser	Ala	Val	Asn	His	Leu	Lys	Ala
65					70				75						80
Asn	Val	Lys	Ser	Ala	Ala	Asp	Leu	Ile	Ser	Leu	Pro	Thr	Thr	Val	Glu
				85					90					95	
Gly	Leu	Gln	Lys	Ser	Val	Ala	Ser	Ile	Gly	Asn	Thr	Leu	Asn	Ser	Val
			100					105					110		
His	Leu	Ala	Val	Glu	Ala	Leu	Gln	Lys	Thr	Val	Asp	Glu	His	Lys	Lys
		115					120					125			
Thr	Met	Glu	Leu	Leu	Gln	Ser	Asp	Met	Asn	Gln	His	Phe	Leu	Lys	Glu
	130					135					140				
Thr	Pro	Gly	Ser	Asn	Gln	Ile	Ile	Pro	Ser	Pro	Ser	Ala	Thr	Ser	Glu
	145				150					155					160
Leu	Asp	Asn	Lys	Thr	His	Ser	Glu	Asn	Leu	Lys	Gln	Met	Gly	Asp	Arg
			165					170						175	
Ser	Ala	Thr	Leu	Lys	Arg	Gln	Ser	Leu	Asp	Gln	Val	Thr	Asn	Arg	Thr
			180					185					190		
Asp	Thr	Val	Lys	Ile	Gln	Lys	Lys	Lys							
			195					200							

<210> 191  
 <211> 379  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -37..-1

<400> 191

Met	Pro	His	Ser	Ser	Leu	His	Pro	Ser	Ile	Pro	Cys	Pro	Arg	Gly	His
	-35						-30				-25				
Gly	Ala	Gln	Lys	Ala	Ala	Leu	Val	Leu	Leu	Ser	Ala	Cys	Leu	Val	Thr
	-20					-15				-10					
Leu	Trp	Gly	Leu	Gly	Glu	Pro	Pro	Glu	His	Thr	Leu	Arg	Tyr	Leu	Val
-5					1				5					10	

Leu	His	Leu	Ala	Ser	Leu	Gln	Leu	Gly	Leu	Leu	Leu	Asn	Gly	Val	Cys
			15					20					25		
Ser	Leu	Ala	Glu	Glu	Leu	Arg	His	Ile	His	Ser	Arg	Tyr	Arg	Gly	Ser
		30					35					40			
Tyr	Trp	Arg	Thr	Val	Arg	Ala	Cys	Leu	Gly	Cys	Pro	Leu	Arg	Arg	Gly
	45					50					55				
Ala	Leu	Leu	Leu	Leu	Ser	Ile	Tyr	Phe	Tyr	Tyr	Ser	Leu	Pro	Asn	Ala
60					65					70				75	
Val	Gly	Pro	Pro	Phe	Thr	Trp	Met	Leu	Ala	Leu	Leu	Gly	Leu	Ser	Gln
				80				85						90	
Ala	Leu	Asn	Ile	Leu	Leu	Gly	Leu	Lys	Gly	Leu	Ala	Pro	Ala	Glu	Ile
		95						100					105		
Ser	Ala	Val	Cys	Glu	Lys	Gly	Asn	Phe	Asn	Val	Ala	His	Gly	Leu	Ala
		110					115					120			
Trp	Ser	Tyr	Tyr	Ile	Gly	Tyr	Leu	Arg	Leu	Ile	Leu	Pro	Glu	Leu	Gln
	125				130						135				
Ala	Arg	Ile	Arg	Thr	Tyr	Asn	Gln	His	Tyr	Asn	Asn	Leu	Leu	Arg	Gly
140					145					150				155	
Ala	Val	Ser	Gln	Arg	Leu	Tyr	Ile	Leu	Leu	Pro	Leu	Asp	Cys	Gly	Val
				160				165						170	
Pro	Asp	Asn	Leu	Ser	Met	Ala	Asp	Pro	Asn	Ile	Arg	Phe	Leu	Asp	Lys
		175					180					185			
Leu	Pro	Gln	Gln	Thr	Gly	Asp	Arg	Ala	Gly	Ile	Lys	Asp	Arg	Val	Tyr
		190					195					200			
Ser	Asn	Ser	Ile	Tyr	Glu	Leu	Leu	Glu	Asn	Gly	Gln	Arg	Ala	Gly	Thr
	205				210						215				
Cys	Val	Leu	Glu	Tyr	Ala	Thr	Pro	Leu	Gln	Thr	Leu	Phe	Ala	Met	Ser
220					225					230				235	
Gln	Tyr	Ser	Gln	Ala	Gly	Phe	Ser	Arg	Glu	Asp	Arg	Leu	Glu	Gln	Ala
				240				245						250	
Lys	Leu	Phe	Cys	Arg	Thr	Leu	Glu	Asp	Ile	Leu	Ala	Asp	Ala	Pro	Glu
		255					260					265			
Ser	Gln	Asn	Asn	Cys	Arg	Leu	Ile	Ala	Tyr	Gln	Glu	Pro	Ala	Asp	Asp
	270						275					280			
Ser	Ser	Phe	Ser	Leu	Ser	Gln	Glu	Val	Leu	Arg	His	Leu	Arg	Gln	Glu
	285				290						295				
Glu	Lys	Glu	Glu	Val	Thr	Val	Gly	Ser	Leu	Lys	Thr	Ser	Ala	Val	Pro
300					305					310				315	
Ser	Thr	Ser	Thr	Met	Ser	Gln	Glu	Pro	Glu	Leu	Leu	Leu	Ser	Gly	Met
				320					325					330	
Gly	Lys	Pro	Leu	Pro	Leu	Arg	Thr	Asp	Phe	Ser					
		335						340							

<210> 192

$\langle 211 \rangle$  112

<212> PRT

<213> Homo sapiens

<400> 192

Met	Pro	Ser	Glu	Gly	Arg	Cys	Trp	Glu	Thr	Leu	Lys	Ala	Leu	Arg	Ser
1				5					10					15	
Ser	Asp	Lys	Gly	Arg	Leu	Cys	Tyr	Tyr	Arg	Asp	Trp	Leu	Leu	Arg	Arg
			20					25					30		
Glu	Asp	Val	Leu	Glu	Glu	Cys	Met	Ser	Leu	Pro	Lys	Leu	Ser	Ser	Tyr
		35					40					45			

```

Ser Gly Trp Val Val Glu His Val Leu Pro His Met Gln Glu Asn Gln
 50          55          60
Pro Leu Ser Glu Thr Ser Pro Ser Ser Thr Ser Ala Ser Ala Leu Asp
65          70          75          80
Gln Pro Ser Phe Val Pro Lys Ser Pro Asp Ala Ser Ser Ala Phe Ser
          85          90          95
Pro Ala Ser Pro Ala Thr Pro Asn Gly Thr Lys Gly Lys Lys Lys Lys
          100          105          110

```

```

<210> 193
<211> 43
<212> PRT
<213> Homo sapiens

```

```

<400> 193
Ser Leu Pro Gln Ala Leu Trp Phe Gln Phe Phe Tyr His Ser Gly Ser
1          5          10          15
Ser Leu Glu Ser Pro Gly Met Leu Asn Gly Pro Phe Gln His Arg Asn
          20          25          30
Ser Arg Ile Met Thr His Arg Ser Ala Glu Lys
          35          40

```

```

<210> 194
<211> 51
<212> PRT
<213> Homo sapiens

```

```

<220>
<221> SIGNAL
<222> -16...-1

```

```

<400> 194
Met Leu Arg Ile Ala Leu Thr Leu Ile Pro Ser Met Leu Ser Arg Ala
-15          -10          -5
Ala Gly Trp Cys Trp Tyr Lys Glu Pro Thr Gln Gln Phe Ser Tyr Leu
1          5          10          15
Cys Leu Pro Cys Leu Ser Trp Asn Lys Lys Gly Asn Val Leu Gln Leu
          20          25          30
Pro Asn Phe
          35

```

```

<210> 195
<211> 244
<212> PRT
<213> Homo sapiens

```

```

<220>
<221> SIGNAL
<222> -18...-1

```

```

<400> 195

```



```

Met Ala Asn Pro Lys Leu Leu Gly Leu Glu Leu Ser Glu Ala Glu Ala
      -15              -10              -5
Ile Gly Ala Asp Ser Ala Arg Phe Glu Glu Leu Leu Leu Gln Ala Ser
      1              5              10
Lys Glu Leu Gln Gln Ala Gln Thr Thr Arg Pro Glu Ser Thr Gln Ile
15              20              25              30
Gln Pro Gln Pro Gly Phe Cys Ile Lys Thr Asn Ser Ser Glu Gly Lys
      35              40              45
Val Phe Ile Asn Ile Cys His Ser Pro Ser Ile Pro Pro Pro Ala Asp
      50              55              60
Val Thr Glu Glu Glu Leu Leu Gln Met Leu Glu Glu Asp Gln Ala Gly
      65              70              75
Phe Arg Ile Pro Met Ser Leu Gly Glu Pro His Ala Glu Leu Asp Ala
      80              85              90
Lys Gly Gln Gly Cys Thr Ala Tyr Asp Val Ala Val Asn Ser Asp Phe
95              100              105              110
Tyr Arg Arg Met Gln Asn Ser Asp Phe Leu Arg Glu Leu Val Ile Thr
      115              120              125
Ile Ala Arg Glu Gly Leu Glu Asp Ile Tyr Asn Leu Gln Leu Asn Pro
      130              135              140
Glu Trp Arg Met Met Lys Asn Arg Pro Phe Met Gly Ser Ile Ser Gln
      145              150              155
Gln Asn Ile Arg Ser Glu Gln Arg Pro Arg Ile Gln Glu Leu Gly Asp
      160              165              170
Leu Tyr Thr Pro Ala Pro Gly Arg Ala Glu Ser Gly Pro Glu Lys Pro
175              180              185              190
His Leu Asn Leu Trp Leu Glu Ala Pro Asp Leu Leu Leu Ala Glu Val
      195              200              205
Asp Leu Pro Lys Leu Asp Gly Ala Leu Gly Leu Ser Leu Glu Ile Gly
      210              215              220
Arg Thr Ala Trp
      225

```

<210> 196  
 <211> 353  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -34..-1

```

<400> 196
Met Glu Arg Gly Leu Lys Ser Ala Asp Pro Arg Asp Gly Thr Gly Tyr
      -30              -25              -20
Thr Gly Trp Ala Gly Ile Ala Val Leu Tyr Leu His Leu Tyr Asp Val
      -15              -10              -5
Phe Gly Asp Pro Ala Tyr Leu Gln Leu Ala His Gly Tyr Val Lys Gln
      1              5              10
Ser Leu Asn Cys Leu Thr Lys Arg Ser Ile Thr Phe Leu Cys Gly Asp
15              20              25              30
Ala Gly Pro Leu Ala Val Ala Ala Val Leu Tyr His Lys Met Asn Asn
      35              40              45
Glu Lys Gln Ala Glu Asp Cys Ile Thr Arg Leu Ile His Leu Asn Lys
      50              55              60

```

```

Ile Asp Pro His Ala Pro Asn Glu Met Leu Tyr Gly Arg Ile Gly Tyr
 65              70              75
Ile Tyr Ala Leu Leu Phe Val Asn Lys Asn Phe Gly Val Glu Lys Thr
 80              85              90
Pro Gln Ser His Ile Gln Gln Ile Cys Glu Thr Ile Leu Thr Ser Gly
 95              100             105             110
Glu Asn Leu Ala Arg Lys Arg Asn Phe Thr Ala Lys Ser Pro Leu Met
              115             120             125
Tyr Glu Trp Tyr Gln Glu Tyr Tyr Val Gly Ala Ala His Gly Leu Ala
              130             135             140
Gly Ile Tyr Tyr Tyr Leu Met Gln Pro Ser Leu Gln Val Ser Gln Gly
              145             150             155
Lys Leu His Ser Leu Val Lys Pro Ser Val Asp Tyr Val Cys Gln Leu
              160             165             170
Lys Phe Pro Ser Gly Asn Tyr Pro Pro Cys Ile Gly Asp Asn Arg Asp
              175             180             185             190
Leu Leu Val His Trp Cys His Gly Ala Pro Gly Val Ile Tyr Met Leu
              195             200             205
Ile Gln Ala Tyr Lys Val Phe Arg Glu Lys Tyr Leu Cys Asp Ala
              210             215             220
Tyr Gln Cys Ala Asp Val Ile Trp Gln Tyr Gly Leu Leu Lys Lys Gly
              225             230             235
Tyr Gly Leu Cys His Gly Ser Ala Gly Asn Ala Tyr Ala Phe Leu Thr
              240             245             250
Leu Tyr Asn Leu Thr Gln Asp Met Lys Tyr Leu Tyr Arg Ala Cys Lys
              255             260             265             270
Phe Ala Glu Trp Cys Leu Glu Tyr Gly Glu His Gly Cys Arg Thr Pro
              275             280             285
Asp Thr Pro Phe Ser Leu Phe Glu Gly Met Ala Gly Thr Ile Tyr Phe
              290             295             300
Leu Ala Asp Leu Leu Val Pro Thr Lys Ala Arg Phe Pro Ala Phe Glu
              305             310             315
Leu

```

<210> 197

<211> 30

<212> PRT

<213> Homo sapiens

<400> 197

```

Met Gln Met Asp Thr Phe Phe Met Ser Glu Lys His Thr His Thr His
1              5              10             15
Thr His Ile His Thr His Thr Arg Lys Thr Lys Lys Lys Lys
              20              25              30

```

<210> 198

<211> 112

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -48..-1

<400> 198

```
Met Gln Asp Thr Gly Ser Val Val Pro Leu His Trp Phe Gly Phe Gly
      -45              -40              -35
Tyr Ala Ala Leu Val Ala Ser Gly Gly Ile Ile Gly Tyr Val Lys Ala
      -30              -25              -20
Gly Ser Val Pro Ser Leu Ala Ala Gly Leu Leu Phe Gly Ser Leu Ala
      -15              -10              -5
Gly Leu Gly Ala Tyr Gln Leu Ser Gln Asp Pro Arg Asn Val Trp Val
1      5      10      15
Phe Leu Ala Thr Ser Gly Thr Leu Ala Gly Ile Met Gly Met Arg Phe
      20      25      30
Tyr His Ser Gly Lys Phe Met Pro Ala Gly Leu Ile Ala Gly Ala Ser
      35      40      45
Leu Leu Met Val Ala Lys Val Gly Val Ser Met Phe Asn Arg Pro His
      50      55      60
```

<210> 199

<211> 54

<212> PRT

<213> Homo sapiens

<400> 199

```
Glu Ile Ala Gly Tyr Gly Ala Glu Gly Phe Ser Ser Val Leu Gly Tyr
      5      10      15
Pro Arg Trp His Arg Leu Pro Pro Gln Ser Leu Gln His His Gln Tyr
      20      25      30
Cys Gln Arg Arg Trp Pro Asp Arg Arg Cys Leu Gln Ser His Thr Gln
      35      40      45
Ser Ser Gly His Leu Pro
      50
```

<210> 200

<211> 151

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -21..-1

<400> 200

```
Met Ala Ala Ser Thr Ser Met Xaa Pro Val Ala Val Thr Ala Ala Val
      -20              -15              -10
Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu Arg Glu Ile
      -5      1      5      10
Lys Lys Gln Leu Leu Ile Ala Gly Leu Thr Arg Glu Arg Gly Leu
      15      20      25
Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser Leu Pro Ala
      30      35      40
Leu Pro Xaa Gly Gln Leu Gln Pro Pro Pro Pro Ile Thr Glu Glu Asp
      45      50      55
Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr Phe Asp Val
```

60		65		70		75									
Lys	Glu	Tyr	Asp	Arg	Ala	Ala	His	Phe	Leu	His	Gly	Cys	Asn	Ser	Lys
			80						85					90	
Lys	Ala	Tyr	Phe	Leu	Tyr	Met	Tyr	Ser	Arg	Tyr	Leu	Val	Arg	Ala	Ile
			95					100					105		
Leu	Lys	Cys	His	Ser	Ala	Phe	Ser	Glu	Thr	Ser	Ile	Phe	Arg	Thr	Asn
		110					115					120			
Gly	Lys	Val	Lys	Ser	Phe	Lys									
	125					130									

<210> 201  
 <211> 228  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -25..-1

<400> 201

Met	Ser	Met	Ala	Val	Glu	Thr	Phe	Gly	Phe	Phe	Met	Ala	Thr	Val	Gly
25					-20				-15						-10
Leu	Leu	Met	Leu	Gly	Val	Thr	Leu	Pro	Asn	Ser	Tyr	Trp	Arg	Val	Ser
			-5					1				5			
Thr	Val	His	Gly	Asn	Val	Ile	Thr	Thr	Asn	Thr	Ile	Phe	Glu	Asn	Leu
	10					15					20				
Trp	Phe	Ser	Cys	Ala	Thr	Asp	Ser	Leu	Gly	Val	Tyr	Asn	Cys	Trp	Glu
25					30						35				
Phe	Pro	Ser	Met	Leu	Ala	Leu	Ser	Gly	Tyr	Ile	Gln	Ala	Cys	Arg	Ala
40				45						50					55
Leu	Met	Ile	Thr	Ala	Ile	Leu	Leu	Gly	Phe	Leu	Gly	Leu	Leu	Leu	Gly
			60					65						70	
Ile	Ala	Gly	Leu	Arg	Cys	Thr	Asn	Ile	Gly	Gly	Leu	Glu	Leu	Ser	Arg
		75					80					85			
Lys	Ala	Lys	Leu	Ala	Ala	Thr	Ala	Gly	Ala	Pro	His	Ile	Leu	Ala	Gly
	90					95						100			
Ile	Cys	Gly	Met	Val	Ala	Ile	Ser	Trp	Tyr	Ala	Phe	Asn	Ile	Thr	Arg
	105					110					115				
Asp	Phe	Phe	Asp	Pro	Leu	Tyr	Pro	Gly	Thr	Lys	Tyr	Glu	Leu	Gly	Pro
120					125					130					135
Ala	Leu	Tyr	Leu	Gly	Trp	Ser	Ala	Ser	Leu	Ile	Ser	Ile	Leu	Gly	Gly
			140					145						150	
Leu	Cys	Leu	Cys	Ser	Ala	Cys	Cys	Cys	Gly	Ser	Asp	Glu	Asp	Pro	Ala
	155						160					165			
Ala	Ser	Ala	Arg	Arg	Pro	Tyr	Gln	Ala	Pro	Val	Ser	Val	Met	Pro	Val
	170					175						180			
Ala	Thr	Ser	Asp	Gln	Glu	Gly	Asp	Ser	Ser	Phe	Gly	Lys	Tyr	Gly	Arg
	185				190						195				
Asn	Ala	Tyr	Val												
200															

<210> 202  
 <211> 64

<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -47..-1

<400> 202  
Met His Gly Phe Glu Ile Ile Ser Leu Lys Glu Glu Ser Pro Leu Gly  
          -45                  -40                  -35  
Lys Val Ser Gln Gly Pro Leu Phe Asn Val Thr Ser Gly Ser Ser Ser  
          -30                  -25                  -20  
Pro Val Thr Trp Leu Gly Leu Leu Ser Phe Gln Asn Leu His Cys Phe  
          -15                  -10                  -5                  1  
Pro Asp Leu Pro Thr Glu Met Pro Leu Arg Ala Lys Gly Val Asn Thr  
              5                          10                          15

<210> 203  
<211> 146  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -31..-1

<400> 203  
Met Met Trp Gln Lys Tyr Ala Gly Ser Arg Arg Ser Met Pro Leu Gly  
          -30                  -25                  -20  
Ala Arg Ile Leu Phe His Gly Val Phe Tyr Ala Gly Gly Phe Ala Ile  
          15                  -10                  -5                  1  
Val Tyr Tyr Leu Ile Gln Lys Phe His Ser Arg Ala Leu Tyr Tyr Lys  
              5                          10                          15  
Leu Ala Val Glu Gln Leu Gln Ser His Pro Glu Ala Gln Glu Ala Leu  
          20                          25                          30  
Gly Pro Pro Leu Asn Ile His Tyr Leu Lys Leu Ile Asp Arg Glu Asn  
          35                          40                          45  
Phe Val Asp Ile Val Asp Ala Lys Leu Lys Ile Pro Val Ser Gly Ser  
          50                          55                          60                          65  
Lys Ser Glu Gly Leu Leu Tyr Val His Ser Ser Arg Gly Gly Pro Phe  
                          70                          75                          80  
Gln Arg Trp His Leu Asp Glu Val Phe Leu Glu Leu Lys Asp Gly Gln  
          85                          90                          95  
Gln Ile Pro Val Phe Lys Leu Ser Gly Glu Asn Gly Asp Glu Val Lys  
          100                          105                          110  
Lys Glu  
          115

<210> 204  
<211> 87  
<212> PRT  
<213> Homo sapiens

<400> 204

```

Met Glu Leu Ala Pro Thr Ala Arg Leu Pro Pro Gly His Gly Ser Leu
1          5          10          15
Pro His Gly Val Leu Gly Pro Arg Ala Thr Gly Ser Val Thr His Leu
          20          25          30
Ser Leu Leu Pro Gln Ile Lys Gln Arg Ala Ser Glu Ala Leu Pro Glu
          35          40          45
Leu Leu Arg Pro Val Thr Pro Ile Thr Asn Phe Glu Gly Ser Gln Ser
          50          55          60
Gln Asp His Ser Gly Ile Phe Gly Leu Val Thr Asn Leu Glu Glu Leu
65          70          75          80
Glu Val Asp Asp Trp Glu Phe
          85

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<210> 205

<211> 40

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -27..-1

<400> 205

```

Met Arg Thr Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr
          -25          -20          -15
Leu Leu Leu Ile Thr Pro Ser Pro Ser Pro Leu Leu Phe Asp Arg Gly
          -10          -5          1          5
Leu Ser Leu Arg Ser Ala Met Ser
          10

```

<210> 206

<211> 154

<212> PRT

<213> Homo sapiens

<400> 206

```

Met Gly Ser Leu Ser Gly Leu Arg Leu Ala Ala Gly Ser Cys Phe Arg
1          5          10          15
Leu Cys Glu Arg Asp Val Ser Ser Ser Leu Arg Leu Thr Arg Ser Ser
          20          25          30
Asp Leu Lys Arg Ile Asn Gly Phe Cys Thr Lys Pro Gln Glu Ser Pro
          35          40          45
Gly Ala Pro Ser Arg Thr Tyr Asn Arg Val Pro Leu His Lys Pro Thr
          50          55          60
Asp Trp Gln Lys Lys Ile Leu Ile Trp Ser Gly Arg Phe Lys Lys Glu
65          70          75          80
Asp Glu Ile Pro Glu Thr Val Ser Leu Glu Met Leu Asp Ala Ala Lys
          85          90          95
Asn Lys Met Arg Val Lys Ser Ser Tyr Leu Met Ile Ala Leu Thr Val
          100          105          110
Val Gly Cys Ile Phe Met Val Ile Glu Gly Lys Lys Ala Ala Gln Arg
          115          120          125

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His Glu Thr Leu Thr Ser Leu Asn Leu Glu Lys Lys Ala Arg Leu Lys  
130 135 140  
Glu Glu Ala Ala Met Lys Ala Lys Thr Glu  
145 150

<210> 207  
<211> 101  
<212> PRT  
<213> Homo sapiens

<400> 207  
Met Val Cys Glu Lys Cys Glu Lys Lys Leu Gly Thr Val Ile Thr Pro  
1 5 10 15  
Asp Thr Trp Lys Asp Gly Ala Arg Asn Thr Thr Glu Ser Gly Gly Arg  
20 25 30  
Lys Leu Asn Lys Asn Lys Ala Leu Thr Ser Lys Lys Ala Arg Phe Asp  
35 40 45  
Pro Tyr Gly Lys Asn Lys Phe Ser Thr Cys Arg Ile Cys Lys Ser Ser  
50 55 60  
Val His Gln Pro Gly Ser His Tyr Cys Gln Gly Cys Ala Tyr Lys Lys  
65 70 75 80  
Gly Ile Cys Ala Met Cys Gly Lys Lys Val Leu Asp Thr Lys Asn Tyr  
85 90 95  
Lys Gln Thr Ser Val  
100

<210> 208  
<211> 456  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -22..-1

<400> 208  
Met Phe Glu Glu Pro Glu Trp Ala Glu Ala Ala Pro Val Ala Ala Gly  
-20 -15 -10  
Leu Gly Pro Val Ile Ser Arg Pro Pro Pro Ala Ala Ser Ser Gln Asn  
-5 1 5 10  
Lys Gly Ser Lys Arg Arg Gln Leu Leu Ala Thr Leu Arg Ala Leu Glu  
15 20 25  
Ala Ala Ser Leu Ser Gln His Pro Pro Ser Leu Cys Ile Ser Asp Ser  
30 35 40  
Glu Glu Glu Glu Glu Arg Lys Lys Lys Cys Pro Lys Lys Ala Ser  
45 50 55  
Phe Ala Ser Ala Ser Ala Glu Val Gly Lys Lys Gly Lys Lys Lys Cys  
60 65 70  
Gln Lys Gln Gly Pro Pro Cys Ser Asp Ser Glu Glu Val Glu Arg  
75 80 85 90  
Lys Lys Lys Cys His Lys Gln Ala Leu Val Gly Ser Asp Ser Ala Glu  
95 100 105  
Asp Glu Lys Arg Lys Arg Lys Cys Gln Lys His Ala Pro Ile Asn Ser

Year	1970	1971	1972	1973	1974	1975	1976	1977	1978	1979	1980	1981	1982	1983	1984	1985	1986	1987	1988	1989	1990	1991	1992	1993	1994	1995	1996	1997	1998	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022	2023	2024	2025	2026	2027	2028	2029	2030	2031	2032	2033	2034	2035	2036	2037	2038	2039	2040	2041	2042	2043	2044	2045	2046	2047	2048	2049	2050	2051	2052	2053	2054	2055	2056	2057	2058	2059	2060	2061	2062	2063	2064	2065	2066	2067	2068	2069	2070	2071	2072	2073	2074	2075	2076	2077	2078	2079	2080	2081	2082	2083	2084	2085	2086	2087	2088	2089	2090	2091	2092	2093	2094	2095	2096	2097	2098	2099	2100
1970	1971	1972	1973	1974	1975	1976	1977	1978	1979	1980	1981	1982	1983	1984	1985	1986	1987	1988	1989	1990	1991	1992	1993	1994	1995	1996	1997	1998	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022	2023	2024	2025	2026	2027	2028	2029	2030	2031	2032	2033	2034	2035	2036	2037	2038	2039	2040	2041	2042	2043	2044	2045	2046	2047	2048	2049	2050	2051	2052	2053	2054	2055	2056	2057	2058	2059	2060	2061	2062	2063	2064	2065	2066	2067	2068	2069	2070	2071	2072	2073	2074	2075	2076	2077	2078	2079	2080	2081	2082	2083	2084	2085	2086	2087	2088	2089	2090	2091	2092	2093	2094	2095	2096	2097	2098	2099	2100	

<211> 98

<213> Homo sapiens

<221> SIGNAL

Met Pro Ser Ser Phe Phe Leu Leu Leu Gln Phe Phe Leu Arg Ile Asp

Gly Val Leu Ile Arg Met Asn Asp Thr Arg Leu Tyr His Glu Ala Asp

Gly Val Leu Ile Arg Met Asn Asp Thr Arg Leu Tyr His Glu Ala Asp



1	5	10	15
Lys Thr Tyr Met	Leu Arg Glu Tyr Thr Ser	Arg Glu Ser Lys	Ile Ser
	20	25	30
Ser Leu Met His	Val Pro Pro Ser	Leu Phe Thr Glu Pro	Asn Glu Ile
	35	40	45
Ser Gln Tyr Leu	Pro Ile Lys Glu	Ala Val Cys Glu	Lys Leu Ile Phe
	50	55	60
Pro Glu Arg Ile	Asp Pro Asn Pro	Ala Asp Ser Gln	Lys Ser Thr Gln
	65	70	75
Val Glu			
80			

<210> 210

<211> 83

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -29..-1

<400> 210

Met Thr Leu Leu	Ser Phe Ala Ala	Phe Thr Ala Ala	Phe Ser Val Leu
	-25	-20	-15
Pro Cys Tyr Tyr	Leu Gly Leu Phe	Gln Arg Ala Leu	Ala Ser Val Phe
	-10	-5	1
Asp Pro Leu Cys	Val Cys Ser Arg	Val Leu Pro Thr	Pro Val Cys Thr
	5	10	15
Leu Val Ala Thr	Gln Ala Glu Lys	Ile Leu Glu Asn	Gly Pro Cys Pro
	20	25	30
Thr Lys Glu Ala	Ala Gln Leu Val	Gly Lys Gly Ser	Val Ser Ala Arg
	40	45	50
Asn Ala Ser			

<210> 211

<211> 229

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -23..-1

<400> 211

Met Gly Asp Lys	Ile Trp Leu Pro	Phe Pro Val Leu	Leu Leu Ala Ala
	-20	-15	-10
Leu Pro Pro Val	Leu Leu Pro Gly	Ala Ala Gly Phe	Thr Pro Ser Leu
	-5	1	5
Asp Ser Asp Phe	Thr Phe Thr Leu	Pro Ala Gly Gln	Lys Glu Cys Phe
	10	15	20
Tyr Gln Pro Met	Pro Leu Lys Ala	Ser Leu Glu Ile	Glu Tyr Gln Val
	30	35	40
Leu Asp Gly Ala	Gly Leu Asp Ile	Asp Phe His Leu	Ala Ser Pro Glu

[illegible] $\langle 210 \rangle \quad 212$  $\langle 211 \rangle$  152

212 PRT

213> Homo sapiens

 $\langle 220 \rangle$ 

<221> SIGNAL

$\langle 222 \rangle$  -21...-1

 $\langle 400 \rangle$  212

Met Ala Gln Leu Gly Ala Val Val Ala Val Ala Ser Ser Phe Phe Cys

-20                      -15                      -10

Ala Ser Leu Phe Ser Ala Val His Lys Ile Glu Glu Gly His Ile Gly

5	1	5	10
1	1	1	1
1	2	2	2
1	3	3	3
1	4	4	4
1	5	5	5
1	6	6	6
1	7	7	7
1	8	8	8
1	9	9	9
1	10	10	10

Val Tyr Tyr Arg Gly Gly Ala Leu Leu Thr Ser Thr Ser Gly Pro Gly

	15	20	25
Plot Width	Meters	Feet	

Phe His Leu Met Leu Pro Phe Ile Thr Ser Tyr Lys Ser Val Gln Thr

30		35		40
Thur Jan Clk	Thur Apr Clk	Hol Jun Apr Hol	Dec 6	al      Fri      al

Thr Leu Gln Thr Asp Glu Val Lys Asn Val Pro Cys Gly Thr Ser Gly  
45 50 55

Gly Val Met Ile Tyr Phe Asn Arg Ile Gly Val Val Asn Phe Leu Val

Gly Val Met Ile Tyr Phe Asp Arg Ile Glu Val Val Asn Phe Leu Val  
60 65 70 75

Pro Asn Ala Val His Asp Ile Val Lys Asp Thr Thr Ala Asp Thr Asp

The Asn His Val His Asp Ile Val Lys Asn Tyr Thr Ala Asp Tyr Asp  
 80 85 90

Lys Ala Leu Ile Phe Asn Lys Ile His His Glu Leu Asn Gln Phe Cys

95 100 105

Ser Val His Thr Leu Gln Glu Val Tyr Ile Glu Leu Phe Gly Leu Glu

110 115 120

Asn Asp Phe Ser Gln Glu Ser Ser

125 130

<210> 213  
 <211> 179  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -54..-1

<400> 213

Met	Ala	Ala	Ser	Glu	Ala	Ala	Val	Val	Ser	Ser	Pro	Ser	Leu	Lys	Thr
				-50					-45					-40	
Asp	Thr	Ser	Pro	Val	Leu	Glu	Thr	Ala	Gly	Thr	Val	Ala	Ala	Met	Ala
			-35					-30					-25		
Ala	Thr	Pro	Ser	Ala	Arg	Ala	Ala	Ala	Val	Val	Ala	Ala	Ala	Ala	Ala
		-20				-15					-10				
Arg	Thr	Gly	Ser	Glu	Ala	Arg	Val	Ser	Lys	Ala	Ala	Leu	Ala	Thr	Lys
	-5				1				5					10	
Leu	Leu	Ser	Leu	Ser	Gly	Val	Phe	Ala	Val	His	Lys	Pro	Lys	Gly	Pro
			15					20					25		
Thr	Ser	Ala	Glu	Leu	Leu	Asn	Arg	Leu	Lys	Glu	Lys	Leu	Leu	Ala	Glu
		30						35					40		
Ala	Gly	Met	Pro	Ser	Pro	Glu	Trp	Thr	Lys	Arg	Lys	Lys	Gln	Thr	Leu
		45				50					55				
Lys	Ile	Gly	His	Gly	Gly	Thr	Leu	Asp	Ser	Ala	Ala	Arg	Gly	Val	Leu
	60				65				70						
Val	Val	Gly	Ile	Gly	Ser	Gly	Thr	Lys	Met	Leu	Thr	Ser	Met	Leu	Ser
	75				80				85					90	
Gly	Ser	Lys	Arg	Tyr	Thr	Ala	Ile	Gly	Glu	Leu	Gly	Lys	Ala	Thr	Asp
			95					100					105		
Thr	Leu	Asp	Ser	Thr	Gly	Lys	Val	Thr	Glu	Glu	Lys	Pro	Tyr	Gly	Met
			110					115					120		
Asn	Leu	Ile													
		125													

<210> 214  
 <211> 269  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -92..-1

<400> 214

Met	Ile	Thr	His	Val	Thr	Leu	Glu	Asp	Ala	Leu	Ser	Asn	Val	Asp	Leu
		-90					-85					-80			
Leu	Glu	Glu	Leu	Pro	Leu	Pro	Asp	Gln	Gln	Pro	Cys	Ile	Glu	Pro	Pro
	-75				-70					-65					
Pro	Ser	Ser	Ile	Met	Tyr	Gln	Ala	Asn	Phe	Asp	Thr	Asn	Phe	Glu	Asp
	-60				-55				-50					-45	
Arg	Asn	Ala	Phe	Val	Thr	Gly	Ile	Ala	Arg	Tyr	Ile	Glu	Gln	Ala	Thr
			-40					-35					-30		
Val	His	Ser	Ser	Met	Asn	Glu	Met	Leu	Glu	Glu	Gly	His	Glu	Tyr	Ala
			-25				-20						-15		

```

Val Met Leu Tyr Thr Trp Arg Ser Cys Ser Arg Ala Ile Pro Gln Val
      -10                -5                1
Lys Cys Asn Glu Gln Pro Asn Arg Val Glu Ile Tyr Glu Lys Thr Val
5      10      15      20
Glu Val Leu Glu Pro Glu Val Thr Lys Leu Met Lys Phe Met Tyr Phe
      25      30      35
Gln Arg Lys Ala Ile Glu Arg Phe Cys Ser Glu Val Lys Arg Leu Cys
      40      45      50
His Ala Glu Arg Arg Lys Asp Phe Val Ser Glu Ala Tyr Leu Leu Thr
      55      60      65
Leu Gly Lys Phe Ile Asn Met Phe Ala Val Leu Asp Glu Leu Lys Asn
      70      75      80
Met Lys Cys Ser Val Lys Asn Asp His Ser Ala Tyr Lys Arg Ala Ala
85      90      95      100
Gln Phe Leu Arg Lys Met Ala Asp Pro Gln Ser Ile Gln Glu Ser Gln
      105      110      115
Asn Leu Ser Met Phe Leu Ala Asn His Asn Arg Ile Thr Gln Cys Leu
      120      125      130
His Gln Gln Leu Glu Val Ile Pro Gly Tyr Glu Glu Leu Leu Ala Asp
      135      140      145
Ile Val Asn Ile Cys Val Asp Tyr Tyr Glu Asn Lys Met Tyr Leu Thr
      150      155      160
Pro Ser Glu Lys His Met Leu Leu Lys Val Lys Leu Pro
165      170      175

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<210> 215

<211> 135

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -22...-1

<400> 215

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Met Gln Thr Val Tyr Tyr Gly Ser Leu Gly Leu Trp Leu Ala Leu Val
      -20                -15                -10
Asp Gly Leu Val Arg Ser Ser Pro Ser Leu Asp Gln Met Phe Asp Ala
      -5      1      5      10
Glu Ile Leu Gly Phe Ser Thr Pro Pro Gly Arg Leu Ser Met Met Ser
      15      20      25
Phe Ile Phe Asn Ala Leu Thr Cys Ala Leu Gly Leu Leu Tyr Phe Ile
      30      35      40
Arg Arg Gly Lys Gln Cys Leu Asp Phe Thr Val Thr Val His Phe Phe
      45      50      55
His Leu Leu Gly Cys Trp Phe Tyr Ser Ser Arg Phe Pro Ser Ala Leu
      60      65      70
Thr Trp Trp Leu Val Gln Ala Val Cys Ile Ala Leu Met Ala Val Ile
      75      80      85      90
Gly Glu Tyr Leu Cys Met Arg Thr Glu Leu Lys Glu Ile Pro Leu Asn
      95      100      105
Ser Ala Pro Lys Ser Asn Val
      110

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<210> 216  
 <211> 67  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -38..-1

<400> 216  
 Met Asn Asn Val Gln Pro Lys Ile Lys His Arg Pro Phe Cys Phe Ser  
                   -35                  -30                  -25  
 Val Lys Gly His Val Lys Met Leu Arg Leu Val Phe Ala Leu Val Thr  
           -20                  -15                  -10  
 Ala Val Cys Cys Leu Ala Asp Gly Ala Leu Ile Tyr Arg Lys Leu Leu  
       -5                  1                  5                  10  
 Phe Asn Pro Asn Gly Pro Tyr Gln Lys Lys Pro Val His Glu Lys Lys  
                   15                  20                  25  
 Glu Val Leu

<210> 217  
 <211> 125  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -54..-1

<400> 217  
 Met Ala Asp Glu Glu Leu Glu Ala Leu Arg Arg Gln Arg Leu Ala Glu  
                   -50                  -45                  -40  
 Leu Gln Ala Lys His Gly Asp Pro Gly Asp Ala Ala Gln Gln Glu Ala  
           -35                  -30                  -25  
 Lys His Arg Glu Ala Glu Met Arg Asn Ser Ile Leu Ala Gln Val Leu  
       -20                  -15                  -10  
 Asp Gln Ser Ala Arg Ala Arg Leu Ser Asn Leu Ala Leu Val Lys Pro  
       -5                  1                  5                  10  
 Glu Lys Thr Lys Ala Val Glu Asn Tyr Leu Ile Gln Met Ala Arg Tyr  
           15                  20                  25  
 Gly Gln Leu Ser Glu Lys Val Ser Glu Gln Gly Leu Ile Glu Ile Leu  
       30                  35                  40  
 Lys Lys Val Ser Gln Gln Thr Glu Lys Thr Thr Thr Val Lys Phe Asn  
       45                  50                  55  
 Arg Arg Lys Val Met Asp Ser Asp Glu Asp Asp Asp Tyr  
       60                  65                  70

<210> 218  
 <211> 376  
 <212> PRT  
 <213> Homo sapiens

<220>

<221> SIGNAL

<222> -21..-1

<400> 218

Met Gly His Arg Phe Leu Arg Gly Leu Leu Thr Leu Leu Leu Pro Pro  
-20 -15 -10  
Pro Pro Leu Tyr Thr Arg His Arg Met Leu Gly Pro Glu Ser Val Pro  
-5 1 5 10  
Pro Pro Lys Arg Ser Arg Ser Lys Leu Met Ala Pro Pro Arg Ile Gly  
15 20 25  
Thr His Asn Gly Thr Phe His Cys Asp Glu Ala Leu Ala Cys Ala Leu  
30 35 40  
Leu Arg Leu Leu Pro Glu Tyr Arg Asp Ala Glu Ile Val Arg Thr Arg  
45 50 55  
Asp Pro Glu Lys Leu Ala Ser Cys Asp Ile Val Val Asp Val Gly Gly  
60 65 70 75  
Glu Tyr Asp Pro Arg Arg His Arg Tyr Asp His His Gln Arg Ser Phe  
80 85 90  
Thr Glu Thr Met Ser Ser Leu Ser Pro Gly Arg Pro Trp Gln Thr Lys  
95 100 105  
Leu Ser Ser Ala Gly Leu Ile Tyr Leu His Phe Gly His Lys Leu Leu  
110 115 120  
Ala Gln Leu Leu Gly Thr Ser Glu Glu Asp Ser Met Val Gly Thr Leu  
125 130 135  
Tyr Asp Lys Met Tyr Glu Asn Phe Val Glu Glu Val Asp Ala Val Asp  
140 145 150 155  
Asn Gly Ile Ser Gln Trp Ala Glu Gly Glu Pro Arg Tyr Ala Leu Thr  
160 165 170  
Thr Thr Leu Ser Ala Arg Val Ala Arg Leu Asn Pro Thr Trp Asn His  
175 180 185  
Pro Asp Gln Asp Thr Glu Ala Gly Phe Lys Arg Ala Met Asp Leu Val  
190 195 200  
Gln Glu Glu Phe Leu Gln Arg Leu Asp Phe Tyr Gln His Ser Trp Leu  
205 210 215  
Pro Ala Arg Ala Leu Val Glu Glu Ala Leu Ala Gln Arg Phe Gln Val  
220 225 230 235  
Asp Pro Ser Gly Glu Ile Val Glu Leu Ala Lys Gly Ala Cys Pro Trp  
240 245 250  
Lys Glu His Leu Tyr His Leu Glu Ser Gly Leu Ser Pro Pro Val Ala  
255 260 265  
Ile Phe Phe Val Ile Tyr Thr Asp Gln Ala Gly Gln Trp Arg Ile Gln  
270 275 280  
Cys Val Pro Lys Glu Pro His Ser Phe Gln Ser Arg Leu Pro Leu Pro  
285 290 295  
Glu Pro Trp Arg Gly Leu Arg Asp Glu Ala Leu Asp Gln Val Ser Gly  
300 305 310 315  
Ile Pro Gly Cys Ile Phe Val His Ala Ser Gly Phe Ile Gly Gly His  
320 325 330  
Arg Thr Arg Glu Gly Ala Leu Ser Met Ala Arg Ala Thr Leu Ala Gln  
335 340 345  
Arg Ser Tyr Leu Pro Gln Ile Ser  
350 355

<210> 219

<211> 211  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -30..-1

<400> 219  
Met Gly Glu Ala Ser Pro Pro Ala Pro Ala Arg Arg His Leu Leu Val  
-30 -25 -20 -15  
Leu Leu Leu Leu Leu Ser Thr Leu Val Ile Pro Ser Ala Ala Ala Pro  
-10 -5 1  
Ile His Asp Ala Asp Ala Gln Glu Ser Ser Leu Gly Leu Thr Gly Leu  
5 10 15  
Gln Ser Leu Leu Gln Gly Phe Ser Arg Leu Phe Leu Lys Gly Asn Leu  
20 25 30  
Leu Arg Gly Ile Asp Ser Leu Phe Ser Ala Pro Met Asp Phe Arg Gly  
35 40 45 50  
Leu Pro Gly Asn Tyr His Lys Glu Glu Asn Gln Glu His Gln Leu Gly  
55 60 65  
Asn Asn Thr Leu Ser Ser His Leu Gln Ile Asp Lys Val Pro Arg Met  
70 75 80  
Glu Glu Lys Glu Ala Leu Val Pro Ile Gln Lys Ala Thr Asp Ser Phe  
85 90 95  
His Thr Glu Leu His Pro Arg Val Ala Phe Trp Ile Ile Lys Leu Pro  
100 105 110  
Arg Arg Arg Ser His Gln Asp Ala Leu Glu Gly Gly His Trp Leu Ser  
115 120 125 130  
Glu Lys Arg His Arg Leu Gln Ala Ile Arg Asp Gly Leu Arg Lys Gly  
135 140 145  
Thr His Lys Asp Val Leu Glu Glu Gly Thr Glu Ser Ser Ser His Ser  
150 155 160  
Arg Leu Ser Pro Arg Lys Thr His Leu Leu Tyr Ile Leu Arg Pro Ser  
165 170 175  
Arg Gln Leu  
180

<210> 220  
<211> 154  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -60..-1

<400> 220  
Met Gly Ser Lys Cys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu  
-60 -55 -50 -45  
Arg Gln Arg Arg Gln Lys Leu Leu Leu Ala Gln Leu His His Arg Lys  
-40 -35 -30  
Arg Val Lys Ala Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu  
-25 -20 -15  
Val Arg Arg Thr Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln

	-10						-5				1								
Cys	Trp	Trp	Arg	Thr	Leu	Val	Gln	Arg	Arg	Ile	Arg	Gln	Arg	Arg	Gln				
5					10					15					20				
Ala	Leu	Leu	Arg	Val	Tyr	Val	Ile	Gln	Glu	Gln	Ala	Thr	Val	Lys	Leu				
				25					30					35					
Gln	Ser	Cys	Ile	Arg	Met	Trp	Gln	Cys	Arg	Gln	Cys	Tyr	Arg	Gln	Met				
			40					45					50						
Cys	Asn	Ala	Leu	Cys	Leu	Phe	Gln	Val	Pro	Glu	Ser	Ser	Leu	Ala	Phe				
	55					60						65							
Gln	Thr	Asp	Gly	Phe	Leu	Gln	Val	Gln	Tyr	Ala	Ile	Pro	Ser	Lys	Gln				
	70				75						80								
Pro	Glu	Phe	His	Ile	Glu	Ile	Leu	Ser	Ile										
85					90														

<210> 221  
 <211> 123  
 <212> PRT  
 <213> Homo sapiens

<220>

<221> SIGNAL

<222> -42..-1

<400> 221

Met	Lys	Gly	Gly	Ala	Phe	Ser	Asn	Leu	Asn	Asp	Ser	Gln	Leu	Ser	Ala				
		-40					-35					-30							
Ser	Phe	Leu	Gln	Pro	Ser	Leu	Gln	Ala	Asn	Cys	Pro	Ala	Leu	Asp	Pro				
	-25					-20					-15								
Ala	Val	Ser	Leu	Ser	Ala	Pro	Ala	Phe	Ala	Ser	Ala	Leu	Arg	Ser	Met				
	-10				-5					1			5						
Lys	Ser	Ser	Gln	Ala	Ala	Arg	Lys	Asp	Asp	Phe	Leu	Arg	Ser	Leu	Ser				
		10					15					20							
Asp	Gly	Asp	Ser	Gly	Thr	Ser	Glu	His	Ile	Ser	Ala	Val	Val	Thr	Ser				
	25						30					35							
Pro	Arg	Ile	Ser	Cys	His	Gly	Ala	Ala	Ile	Pro	Thr	Ala	Arg	Ala	Leu				
	40					45					50								
Cys	Leu	Gly	Cys	Ser	Cys	Cys	Thr	Glu	Arg	Leu	Leu	Leu	Pro	Pro	Pro				
	55				60					65					70				
Ser	Leu	Leu	Ser	Leu	Glu	Ala	Pro	Ala	Ser	Thr									
				75					80										

<210> 222  
 <211> 346  
 <212> PRT  
 <213> Homo sapiens

<220>

<221> SIGNAL

<222> -19..-1

<400> 222

Met	Ala	Met	Ala	Gln	Lys	Leu	Ser	His	Leu	Leu	Pro	Ser	Leu	Arg	Gln				
				-15					-10					-5					



Val	Ile	Gln	Glu	Pro	Gln	Leu	Ser	Leu	Gln	Pro	Glu	Pro	Val	Phe	Thr
		1				5				10					
Val	Asp	Arg	Ala	Glu	Val	Pro	Pro	Leu	Phe	Trp	Lys	Pro	Tyr	Ile	Tyr
15						20				25					
Ala	Gly	Tyr	Arg	Pro	Leu	His	Gln	Thr	Trp	Arg	Phe	Tyr	Phe	Arg	Thr
30					35				40					45	
Leu	Phe	Gln	Gln	His	Asn	Glu	Ala	Val	Asn	Val	Trp	Thr	His	Leu	Leu
				50					55					60	
Ala	Ala	Leu	Val	Leu	Leu	Leu	Arg	Leu	Ala	Leu	Phe	Val	Glu	Thr	Val
			65				70					75			
Asp	Phe	Trp	Gly	Asp	Pro	His	Ala	Leu	Pro	Leu	Phe	Ile	Ile	Val	Leu
80						85					90				
Ala	Ser	Phe	Thr	Tyr	Leu	Ser	Leu	Ser	Ala	Leu	Ala	His	Leu	Leu	Gln
95						100					105				
Ala	Lys	Ser	Glu	Phe	Trp	His	Tyr	Ser	Phe	Phe	Phe	Leu	Asp	Tyr	Val
110					115					120				125	
Gly	Val	Ala	Val	Tyr	Gln	Phe	Gly	Ser	Ala	Leu	Ala	His	Phe	Tyr	Tyr
				130					135					140	
Ala	Ile	Glu	Pro	Ala	Trp	His	Ala	Gln	Val	Gln	Ala	Val	Phe	Leu	Pro
			145				150					155			
Met	Ala	Ala	Phe	Leu	Ala	Trp	Leu	Ser	Cys	Ile	Gly	Ser	Cys	Tyr	Asn
		160				165					170				
Lys	Tyr	Ile	Gln	Lys	Pro	Gly	Leu	Leu	Gly	Arg	Thr	Cys	Gln	Glu	Val
175						180				185					
Pro	Ser	Val	Leu	Ala	Tyr	Ala	Leu	Asp	Ile	Ser	Pro	Val	Val	His	Arg
190					195				200					205	
Ile	Phe	Val	Ser	Ser	Asp	Pro	Thr	Thr	Asp	Asp	Pro	Ala	Leu	Leu	Tyr
			210						215					220	
His	Lys	Cys	Gln	Val	Val	Phe	Phe	Leu	Leu	Ala	Ala	Ala	Phe	Phe	Ser
			225				230					235			
Thr	Phe	Met	Pro	Glu	Arg	Trp	Phe	Pro	Gly	Ser	Cys	His	Val	Phe	Gly
		240				245					250				
Gln	Gly	His	Gln	Leu	Phe	His	Ile	Phe	Leu	Val	Leu	Cys	Thr	Leu	Ala
		255				260					265				
Gln	Leu	Glu	Ala	Val	Ala	Leu	Asp	Tyr	Glu	Ala	Arg	Arg	Pro	Ile	Tyr
270					275					280				285	
Glu	Pro	Leu	His	Thr	His	Trp	Pro	His	Asn	Phe	Ser	Gly	Leu	Phe	Leu
			290						295					300	
Leu	Thr	Val	Gly	Ser	Ser	Ile	Leu	Thr	Ala	Phe	Leu	Leu	Ser	Gln	Leu
			305						310					315	
Val	Gln	Arg	Lys	Leu	Asp	Gln	Lys	Thr	Lys						
		320					325								

<210> 223  
 <211> 210  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -20...-1

<400> 223  
 Met Asp Asn Arg Phe Ala Thr Ala Phe Val Ile Ala Cys Val Leu Ser  
 -20 -15 -10 -5

```

Leu Ile Ser Thr Ile Tyr Met Ala Ala Ser Ile Gly Thr Asp Phe Trp
      1              5              10
Tyr Glu Tyr Arg Ser Pro Val Gln Glu Asn Ser Ser Asp Leu Asn Lys
      15              20              25
Ser Ile Trp Asp Glu Phe Ile Ser Asp Glu Ala Asp Glu Lys Thr Tyr
      30              35              40
Asn Asp Ala Leu Phe Arg Tyr Asn Gly Thr Val Gly Leu Trp Arg Arg
      45              50              55              60
Cys Ile Thr Ile Pro Lys Asn Met His Trp Tyr Ser Pro Pro Glu Arg
      65              70              75
Thr Glu Ser Phe Asp Val Val Thr Lys Cys Val Ser Phe Thr Leu Thr
      80              85              90
Glu Gln Phe Met Glu Lys Phe Val Asp Pro Gly Asn His Asn Ser Gly
      95              100              105
Ile Asp Leu Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu Leu Pro
      110              115              120
Phe Val Ser Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly Leu Cys
      125              130              135              140
Ala Cys Ile Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly Ile Leu
      145              150              155
His Leu Leu Ala Val Thr Lys Glu Ser Met Leu Pro Ala Gly Ala Glu
      160              165              170
Ser Lys His Thr Ala Thr Pro Ala His Ala Cys Val Gln Thr Gly Lys
      175              180              185
Pro Lys
      190

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<210> 224

<211> 184

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -20...-1

<400> 224

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Met Asp Asn Arg Phe Ala Thr Ala Phe Val Ile Ala Cys Val Leu Ser
-20              -15              -10              -5
Leu Ile Ser Thr Ile Tyr Met Ala Ala Ser Ile Gly Thr Asp Phe Trp
      1              5              10
Tyr Glu Tyr Arg Ser Pro Val Gln Glu Asn Ser Ser Asp Leu Asn Lys
      15              20              25
Ser Ile Trp Asp Glu Phe Ile Ser Asp Glu Ala Asp Glu Lys Thr Tyr
      30              35              40
Asn Asp Ala Pro Phe Arg Tyr Asn Gly Thr Val Gly Leu Trp Arg Arg
      45              50              55              60
Cys Ile Thr Ile Pro Lys Asn Met His Trp Tyr Ser Pro Pro Glu Arg
      65              70              75
Thr Glu Ser Phe Asp Val Val Thr Lys Cys Val Ser Phe Thr Leu Thr
      80              85              90
Glu Gln Phe Met Glu Lys Phe Val Asp Pro Gly Asn His Asn Ser Gly
      95              100              105
Ile Asp Leu Leu Arg Thr Tyr Leu Trp Arg Cys Gln Phe Leu Leu Pro
      110              115              120

```

Phe Val Ser Leu Gly Leu Met Cys Phe Gly Ala Leu Ile Gly Leu Cys  
 125 130 135 140  
 Ala Cys Ile Cys Arg Ser Leu Tyr Pro Thr Ile Ala Thr Gly Ile Leu  
 145 150 155  
 His Leu Leu Ala Asp Thr Met Leu  
 160

<210> 225  
 <211> 227  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -22..-1

<400> 225

Met Gly Trp Thr Met Arg Leu Val Thr Ala Ala Leu Leu Leu Gly Leu  
 -20 -15 -10  
 Met Met Val Val Thr Gly Asp Glu Asp Glu Asn Ser Pro Cys Ala His  
 -5 1 5 10  
 Glu Ala Leu Leu Asp Glu Asp Thr Leu Phe Cys Gln Gly Leu Glu Val  
 15 20 25  
 Phe Tyr Pro Glu Leu Gly Asn Ile Gly Cys Lys Val Val Pro Asp Cys  
 30 35 40  
 Asn Asn Tyr Arg Gln Lys Ile Thr Ser Trp Met Glu Pro Ile Val Lys  
 45 50 55  
 Phe Pro Gly Ala Val Asp Gly Ala Thr Tyr Ile Leu Val Met Val Asp  
 60 65 70  
 Pro Asp Ala Pro Ser Arg Ala Glu Pro Arg Gln Arg Phe Trp Arg His  
 75 80 85 90  
 Trp Leu Val Thr Asp Ile Lys Gly Ala Asp Leu Lys Lys Gly Lys Ile  
 95 100 105  
 Gln Gly Gln Glu Leu Ser Ala Tyr Gln Ala Pro Ser Pro Pro Ala His  
 110 115 120  
 Ser Gly Phe His Arg Tyr Gln Phe Phe Val Tyr Leu Gln Glu Gly Lys  
 125 130 135  
 Val Ile Ser Leu Leu Pro Lys Glu Asn Lys Thr Arg Gly Ser Trp Lys  
 140 145 150  
 Met Asp Arg Phe Leu Asn Arg Phe His Leu Gly Glu Pro Glu Ala Ser  
 155 160 165 170  
 Thr Gln Phe Met Thr Gln Asn Tyr Gln Asp Ser Pro Thr Leu Gln Ala  
 175 180 185  
 Pro Arg Glu Arg Ala Ser Glu Pro Lys His Lys Asn Gln Ala Glu Ile  
 190 195 200  
 Ala Ala Cys  
 205

<210> 226  
 <211> 74  
 <212> PRT  
 <213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -41..-1

<400> 226  
Met Ile Ala Arg Arg Asn Pro Val Pro Leu Arg Phe Leu Pro Asp Glu  
-40 -35 -30  
Ala Arg Ser Leu Pro Pro Pro Lys Leu Thr Asp Pro Arg Leu Leu Tyr  
-25 -20 -15 -10  
Ile Gly Phe Leu Gly Tyr Cys Ser Gly Leu Ile Asp Asn Leu Ile Arg  
-5 1 5  
Arg Arg Pro Ile Ala Thr Ala Gly Leu His Arg Gln Leu Leu Tyr Ile  
10 15 20  
Thr Ala Phe Phe Leu Leu Asp Ile Ile Leu  
25 30

<210> 227  
<211> 73  
<212> PRT  
<213> Homo sapiens

<400> 227  
Met Glu Lys Tyr Glu Asn Leu Gly Leu Val Gly Glu Gly Ser Tyr Gly  
5 10 15  
Met Val Met Lys Cys Arg Asn Lys Asp Thr Gly Arg Ile Val Ala Ile  
20 25 30  
Lys Lys Phe Leu Glu Ser Asp Asp Asp Lys Met Val Lys Lys Ile Ala  
35 40 45  
Met Arg Glu Val Lys Leu Leu Lys Gln Leu Arg His Glu Asn Leu Val  
50 55 60  
Asn Leu Leu Glu Val Cys Lys Lys Lys  
65 70

<210> 228  
<211> 82  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -16..-1

<400> 228  
Met Lys Arg Leu Leu Pro Ala Thr Ser Leu Ala Gly Pro Val Leu Ser  
-15 -10 -5  
Thr Leu Ile Ala Pro Thr Pro Met Leu Phe Cys Glu Asp Lys Ser Trp  
1 5 10 15  
Asp Leu Phe Leu Phe Phe Lys Ser His Lys Thr Trp Gly Ile Ser Thr  
20 25 30  
Asn Leu Ser Ser Cys Pro Phe Gly Asn Leu Phe Leu Cys Val Gln Phe  
35 40 45  
Val Arg Glu Lys Gln Ser Phe Cys Met Asn Thr Glu Cys Asp Leu Arg  
50 55 60

Lys Asn  
65

<210> 229  
<211> 119  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -56...-1

<400> 229  
Met Ala Glu Pro Ser Ala Ala Thr Gln Ser His Ser Ile Ser Ser Ser  
-55 -50 -45  
Ser Phe Gly Ala Glu Pro Ser Ala Pro Gly Gly Gly Gly Ser Pro Gly  
-40 -35 -30 -25  
Ala Cys Pro Ala Leu Gly Thr Lys Ser Cys Ser Ser Ser Cys Ala Asp  
-20 -15 -10  
Ser Phe Val Ser Ser Ser Ser Ser Gln Pro Val Ser Leu Phe Ser Thr  
-5 1 5  
Ser Gln Glu Gly Leu Ser Ser Leu Cys Ser Asp Glu Pro Ser Ser Glu  
10 15 20  
Ile Met Thr Ser Ser Phe Leu Ser Ser Ser Glu Ile His Asn Thr Gly  
25 30 35 40  
Leu Thr Ile Leu His Gly Glu Lys Ser His Val Leu Gly Ser Gln Pro  
45 50 55  
Ile Leu Ala Lys Lys Lys  
60

<210> 230  
<211> 54  
<212> PRT  
<213> Homo sapiens

<400> 230  
Ala Phe Val Trp Glu Pro Ala Met Val Arg Ile Asn Ala Leu Thr Ala  
1 5 10 15  
Ala Ser Glu Ala Ala Cys Leu Ile Val Ser Val Asp Glu Thr Ile Lys  
20 25 30  
Asn Pro Arg Ser Thr Val Asp Ala Pro Thr Ala Ala Gly Arg Gly Arg  
35 40 45  
Gly Arg Gly Arg Pro His  
50

<210> 231  
<211> 210  
<212> PRT  
<213> Homo sapiens

<220>

<221> SIGNAL  
<222> -14...-1

<400> 231

Met Leu Thr Leu Leu Gly Leu Ser Phe Ile Leu Ala Gly Leu Ile Val  
                  -10                  -5                  1  
Gly Gly Ala Cys Ile Tyr Lys Tyr Phe Met Pro Lys Ser Thr Ile Tyr  
          5                  10                  15  
Arg Gly Glu Met Cys Phe Phe Asp Ser Glu Asp Pro Ala Asn Ser Leu  
          20                  25                  30  
Arg Gly Gly Glu Pro Asn Phe Leu Pro Val Thr Glu Glu Ala Asp Ile  
35                  40                  45                  50  
Arg Glu Asp Asp Asn Ile Ala Ile Ile Asp Val Pro Val Pro Ser Phe  
                  55                  60                  65  
Ser Asp Ser Asp Pro Ala Ala Ile Ile His Asp Phe Glu Lys Gly Met  
          70                  75                  80  
Thr Ala Tyr Leu Asp Leu Leu Leu Gly Ile Cys Tyr Leu Met Pro Leu  
          85                  90                  95  
Asn Thr Ser Ile Val Met Pro Pro Lys Asn Leu Val Glu Leu Phe Gly  
100                  105                  110  
Lys Leu Ala Ser Gly Arg Tyr Leu Pro Gln Thr Tyr Val Val Arg Glu  
115                  120                  125                  130  
Asp Leu Val Ala Val Glu Glu Ile Arg Asp Val Ser Asn Leu Gly Ile  
                  135                  140                  145  
Phe Ile Tyr Gln Leu Cys Asn Asn Arg Lys Ser Phe Arg Leu Arg Arg  
                  150                  155                  160  
Arg Asp Leu Leu Leu Gly Phe Asn Lys Arg Ala Ile Asp Lys Cys Trp  
          165                  170                  175  
Lys Ile Arg His Phe Pro Asn Glu Phe Ile Val Glu Thr Lys Ile Cys  
180                  185                  190  
Gln Glu  
195

<210> 232  
<211> 108  
<212> PRT

<213> Homo sapiens

<400> 232

Met Gly Cys Val Phe Gln Ser Thr Glu Asp Lys Cys Ile Phe Lys Ile  
1                  5                  10                  15  
Asp Trp Thr Leu Ser Pro Gly Glu His Ala Lys Asp Glu Tyr Val Leu  
          20                  25                  30  
Tyr Tyr Tyr Ser Asn Leu Ser Val Pro Ile Gly Arg Phe Gln Asn Arg  
          35                  40                  45  
Val His Leu Met Gly Asp Ile Leu Cys Asn Asp Gly Ser Leu Leu Leu  
50                  55                  60  
Gln Asp Val Gln Glu Ala Asp Gln Gly Thr Tyr Ile Cys Glu Ile Arg  
65                  70                  75                  80  
Leu Lys Gly Glu Ser Gln Val Phe Lys Lys Ala Val Val Leu His Val  
          85                  90                  95  
Leu Pro Glu Glu Pro Lys Gly Thr Gln Met Leu Thr  
100                  105

<210> 233  
 <211> 43  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -18..-1

<400> 233  
 Met Ser Ser Gly Arg Leu Arg Trp Leu Met Pro Val Ile Pro Ala Leu  
                   -15                  -10                  -5  
 Trp Gly Ala Glu Lys Gly Glu Ser Pro Glu Val Ser Ser Phe Glu Thr  
           1                  5                  10  
 Arg Leu Ala Asn Met Ala Lys Pro Cys Leu Tyr  
 15                  20                  25

<210> 234  
 <211> 36  
 <212> PRT  
 <213> Homo sapiens

<400> 234  
 Met Ser Ala Arg Ile Pro Phe Tyr Lys Asp Thr Ser Gln Ile Arg Leu  
                   5                  10                  15  
 Gly Ser Thr Ile Ile Pro His Phe Asn Leu Ile Thr Phe Val Lys Thr  
                   20                  25                  30  
 Phe Phe Gln Ile  
                   35

<210> 235  
 <211> 307  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -13..-1

<400> 235  
 Met Leu Ala Val Ser Leu Thr Val Pro Leu Leu Gly Ala Met Met Leu  
                   -10                  -5                  1  
 Leu Glu Ser Pro Ile Asp Pro Gln Pro Leu Ser Phe Lys Glu Pro Pro  
           5                  10                  15  
 Leu Leu Leu Gly Val Leu His Pro Asn Thr Lys Leu Arg Gln Ala Glu  
 20                  25                  30                  35  
 Arg Leu Phe Glu Asn Gln Leu Val Gly Pro Glu Ser Ile Ala His Ile  
                   40                  45                  50  
 Gly Asp Val Met Phe Thr Gly Thr Ala Asp Gly Arg Val Val Lys Leu  
           55                  60                  65  
 Glu Asn Gly Glu Ile Glu Thr Ile Ala Arg Phe Gly Ser Gly Pro Cys  
           70                  75                  80

[illegible]

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<210> 236
<211> 106
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -32..-1

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<400> 236															
Met	Phe	Ala	Pro	Ala	Val	Met	Arg	Ala	Phe	Arg	Lys	Asn	Lys	Thr	Leu
		-30					-25					-20			
Gly	Tyr	Gly	Val	Pro	Met	Leu	Leu	Leu	Ile	Val	Gly	Gly	Ser	Phe	Gly
		-15				-10					-5				
Leu	Arg	Glu	Phe	Ser	Gln	Ile	Arg	Tyr	Asp	Ala	Val	Lys	Ser	Lys	Met
1				5					10					15	
Asp	Pro	Glu	Leu	Glu	Lys	Lys	Leu	Lys	Glu	Asn	Lys	Ile	Ser	Leu	Glu
			20					25					30		
Ser	Glu	Tyr	Glu	Lys	Ile	Lys	Asp	Ser	Lys	Phe	Asp	Asp	Trp	Lys	Asn
		35					40					45			
Ile	Arg	Gly	Pro	Arg	Pro	Trp	Glu	Asp	Pro	Asp	Leu	Leu	Gln	Gly	Arg
		50				55					60				
Asn	Pro	Glu	Ser	Leu	Lys	Thr	Lys	Thr	Thr						
65					70										



<210> 237  
<211> 42  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -19..-1

<400> 237  
Met Asp Leu Arg Gln Phe Leu Met Cys Leu Ser Leu Cys Thr Ala Phe  
                  -15                  -10                  -5  
Ala Leu Ser Lys Pro Thr Glu Lys Lys Asp Arg Val His His Glu Pro  
                  1                  5                  10  
Gln Leu Ser Asp Lys Val His Asn Asp Ile  
          15                  20

<210> 238  
<211> 117  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -20..-1

<400> 238  
Met Asp Asn Arg Phe Ala Thr Ala Phe Val Ile Ala Cys Val Leu Ser  
-20                  -15                  -10                  -5  
Leu Ile Ser Thr Ile Tyr Met Ala Ala Ser Ile Gly Thr Asp Phe Trp  
                  1                  5                  10  
Tyr Glu Tyr Arg Ser Pro Val Gln Glu Asn Ser Ser Asp Leu Asn Lys  
          15                  20                  25  
Ser Ile Trp Asp Glu Phe Ile Ser Asp Glu Ala Asp Glu Lys Thr Tyr  
          30                  35                  40  
Asn Asp Ala Leu Phe Arg Tyr Asn Gly Thr Val Gly Leu Trp Gly Arg  
          45                  50                  55                  60  
Cys Ile Thr Ile Pro Lys Asn Met His Trp Tyr Ser Pro Pro Glu Arg  
                  65                  70                  75  
Thr Gly Ile Ser Leu Ile Leu Thr Ser Val Phe Phe Thr Trp Leu Ile  
          80                  85                  90  
Ile Asp Lys Thr Thr  
          95

<210> 239  
<211> 178  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -37..-1

<400> 239

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Met Glu Arg Gln Ser Arg Val Met Ser Glu Lys Asp Glu Tyr Gln Phe
  -35                      -30                      -25
Gln His Xaa Xaa Ala Xaa Xaa Leu Leu Val Phe Asn Phe Leu Leu Ile
  -20                      -15                      -10
Leu Thr Ile Leu Thr Ile Trp Leu Phe Lys Asn His Arg Phe Arg Phe
  -5                      1                      5                      10
Leu His Glu Thr Gly Gly Ala Met Val Tyr Gly Leu Ile Met Gly Leu
      15                      20                      25
Ile Ser Arg Tyr Ala Thr Ala Pro Thr Asp Ile Glu Ser Gly Thr Val
      30                      35                      40
Cys Asp Cys Val Lys Leu Thr Phe Ser Pro Pro Thr Leu Leu Val Asn
      45                      50                      55
Val Thr Asp Gln Val Tyr Glu Tyr Lys Tyr Lys Arg Glu Ile Ser Gln
      60                      65                      70                      75
His Asn Ile Asn Pro His Gln Gly Asn Ala Ile Leu Glu Lys Met Thr
      80                      85                      90
Phe Asp Pro Glu Ile Phe Phe Asn Val Leu Leu Pro Pro Ile Ile Phe
      95                      100                      105
His Ala Gly Tyr Ser Leu Lys Lys Arg His Phe Phe Gln Asn Leu Gly
      110                      115                      120
Ser Ile Leu Thr Tyr Ala Phe Leu Gly Thr Ala Ile Ser Cys Ile Val
      125                      130                      135
Ile Gly
140

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<210> 240

<211> 126

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -27..-1

<400> 240

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Met Gln Phe Val Asn Val Gly Tyr Phe Leu Ile Ala Ala Gly Val Val
  -25                      -20                      -15
Val Leu Ala Leu Gly Phe Leu Gly Cys Tyr Gly Ala Lys Thr Glu Ser
  -10                      -5                      1                      5
Met Cys Ala Leu Val Thr Phe Phe Phe Ile Leu Leu Leu Ile Phe Ile
      10                      15                      20
Ala Glu Val Ala Ala Ala Val Val Ala Leu Val Tyr Thr Thr Met Ala
      25                      30                      35
Glu His Phe Leu Thr Leu Leu Val Val Pro Ala Ile Lys Lys Asp Tyr
      40                      45                      50
Gly Ser Gln Glu Asp Phe Thr Gln Val Trp Asn Thr Thr Met Lys Gly
      55                      60                      65
Leu Lys Cys Arg Gly Phe Thr Asn Tyr Thr Asp Phe Glu Asp Ser Pro
      70                      75                      80                      85
Tyr Phe Lys Met His Lys Pro Val Thr Met Lys Lys Lys Lys
      90                      95

```

<210> 241  
<211> 174  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -115..-1

<400> 241  
Met Arg Trp Ser Cys Glu His Leu Val Met Val Trp Ile Asn Ala Phe  
-115 -110 -105 -100  
Val Met Leu Thr Thr Gln Leu Leu Pro Ser Lys Tyr Cys Asp Leu Leu  
-95 -90 -85  
His Lys Ser Ala Ala His Leu Gly Lys Trp Gln Lys Leu Glu His Gly  
-80 -75 -70  
Ser Tyr Ser Asn Ala Pro Gln His Ile Trp Ser Glu Asn Thr Ile Trp  
-65 -60 -55  
Pro Gln Gly Val Leu Val Arg His Ser Arg Cys Leu Tyr Arg Ala Met  
-50 -45 -40  
Gly Pro Tyr Asn Val Ala Val Pro Ser Asp Val Ser His Ala Arg Phe  
35 -30 -25 -20  
Tyr Phe Leu Phe His Arg Pro Leu Arg Leu Leu Asn Leu Leu Ile Leu  
-15 -10 -5  
Ile Glu Gly Gly Val Val Phe Tyr Gln Leu Tyr Ser Leu Leu Arg Ser  
1 5 10  
Glu Lys Trp Asn His Thr Leu Ser Met Ala Leu Ile Leu Phe Cys Asn  
15 20 25  
Tyr Tyr Val Leu Phe Lys Leu Leu Arg Asp Arg Ile Val Leu Gly Arg  
30 35 40 45  
Ala Tyr Ser Tyr Pro Leu Asn Ser Tyr Glu Leu Lys Ala Asn  
50 55

<210> 242  
<211> 896  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 18..173

<221> sig\_peptide  
<222> 18..77  
<223> Von Heijne matrix  
score 6.5  
seq GLCVLQLTTAVTS/AF

<221> polyA\_signal  
<222> 864..869

<221> polyA\_site  
<222> 882..893

<400> 242

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aaccttcaca gtgtgag atg cct agt gtg aac agt gct gga tta tgt gtc      50
              Met Pro Ser Val Asn Ser Ala Gly Leu Cys Val
              -20              -15              -10
ttg cag ttg aca acg gca gtr acc agt gcc ttt tta cta gca aaa gtg      98
Leu Gln Leu Thr Thr Ala Val Thr Ser Ala Phe Leu Leu Ala Lys Val
              -5              1              5
aat cct ttc gaa rct ttt ctc tca agg ggc ttt tgg cta tgt gct gcc      146
Asn Pro Phe Glu Xaa Phe Leu Ser Arg Gly Phe Trp Leu Cys Ala Ala
              10              15              20
cat cat ttc att cat cct tgc ctg gat tgagacgtgt tcctgattca      193
His His Phe Ile His Pro Cys Leu Asp
              25              30
aagtgttacc tcaagaagca gaagaagaaa acagactcct gatagttcag gatgcttcag      253
agagggcagc acttatacct ggtgggtcttt ctgatgggtca gttttattcc ctcctgaat      313
ccgaagcagg atctgaagaa gctgaagaaa aacaggacag tgagaaacca cttttagaac      373
tatgagtact acttttgtaa aatgtgaaaa accctcacag aaagtcacg aggcaaaaag      433
aggcaggcag tggagtctcc ctgtcgacag taaagttgaa atggtgacgt ccaactgctgg      493
ctttattgaa cagctaataa agatttattt attgtaatac ctcacagacg ttgtaccata      553
tccatgcaca tttagttgcc tgcctgtggc tggtaaggta atgtcatgat tcctcctctc      613
ttcagtgaga ctgagcctga tgtgttaaca aataggtgaa gaaagtcttg tgctgtattc      673
ctaatacaaaa gacttaatat attgaagtaa cactttttta gtaagcaaga taccttttta      733
tttcaattca cagaatggaa tttttttgtt tcatgtctca gatttatttt gtatttcctt      793
tttaacactc tacatttccc ttgtttttta actcatgcac atgtgctctt tgtacagttt      853
caaaaagtgt aataaaatct gacatgtcaa araaaaaaa mcY      896

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<210> 243

<211> 851

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 17..595

<221> sig\_peptide

<222> 17..85

<223> Von Heijne matrix

score 3.70000004768372

seq FLPLPLXRAFCRG/CQ

<221> polyA\_signal

<222> 820..825

<221> polyA\_site

<222> 840..851

<400> 243

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aagggggcgt ggggcc atg gtg gtc ttg cgg gcg ggg aag aag acc ttt ctc      52
              Met Val Val Leu Arg Ala Gly Lys Lys Thr Phe Leu
              -20              -15
ccc cct ctm wgc cgc gcc ttc gcc tgc cgc ggc tgt caa ctc gct ccg      100
Pro Pro Leu Xaa Arg Ala Phe Ala Cys Arg Gly Cys Gln Leu Ala Pro
              -10              -5              1              5
gag cgc ggc gcc gag cgc agg gat aca gcg ccc agc ggg gtc tca aga      148
Glu Arg Gly Ala Glu Arg Arg Asp Thr Ala Pro Ser Gly Val Ser Arg

```

```

      10      15      20
ttc tgc cct cca aga aag tct tgc cat gat tgg ata gga ccc cca gat 196
Phe Cys Pro Pro Arg Lys Ser Cys His Asp Trp Ile Gly Pro Pro Asp
      25      30      35
aaa tat tca aac ctt cga cct gtt cac ttt tac ata cct gaa aat gaa 244
Lys Tyr Ser Asn Leu Arg Pro Val His Phe Tyr Ile Pro Glu Asn Glu
      40      45      50
tct cca ttg gaa caa aag ctt aga aaa tta aga caa gaa aca caa gaa 292
Ser Pro Leu Glu Gln Lys Leu Arg Lys Leu Arg Gln Glu Thr Gln Glu
      55      60      65
tgg aat caa cag ttc tgg gca aac cag aat ttg act ttt agt aag gaa 340
Trp Asn Gln Gln Phe Trp Ala Asn Gln Asn Leu Thr Phe Ser Lys Glu
      70      75      80      85
aaa gaa gaa ttt att cac tca aga cta aaa act aaa ggc ctg ggc ctg 388
Lys Glu Glu Phe Ile His Ser Arg Leu Lys Thr Lys Gly Leu Gly Leu
      90      95      100
aga act gaa tca ggt cag aaa gca aca ttg aat gca gaa gaa atg gcg 436
Arg Thr Glu Ser Gly Gln Lys Ala Thr Leu Asn Ala Glu Glu Met Ala
      105      110      115
gac ttc tac aag gaa ttt tta agt aaa aat ttt cag aag cac atg tat 484
Asp Phe Tyr Lys Glu Phe Leu Ser Lys Asn Phe Gln Lys His Met Tyr
      120      125      130
tat aac aga gat tgg tac aag cgc aat ttt gcc atc acc ttc ttc atg 532
Tyr Asn Arg Asp Trp Tyr Lys Arg Asn Phe Ala Ile Thr Phe Phe Met
      135      140      145
gga aaa gtg gcc ctg gaa agg att tgg aac aag ctt aaa cag aaa caa 580
Gly Lys Val Ala Leu Glu Arg Ile Trp Asn Lys Leu Lys Gln Lys Gln
      150      155      160      165
aag aag agg agc aac taggagtcga ctctgaccga gccagagtcg aggtttccac 635
Lys Lys Arg Ser Asn
      170
aggaagcara tggagctcct ttcacagggg ctctgagaaa aactggagct gatctcaaga 695
agccccacat cttcctaagg ggccccatgg cctgtttggg ggcagggtag gtcctggggc 755
actgtggggc gcctgcctgc tgatgtgggc tctaggccag cttgttgtca cgtacgtggg 815
gtgaaataaa gcccaagcac tgggaaaaaa aaaaaa 851

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<210> 244

<211> 495

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 89..334

<221> sig\_peptide

<222> 89..130

<223> Von Heijne matrix

score 3.59999990463257

seq AFTLXSLQLQAALL/CV

<221> polyA\_signal

<222> 462..467

<221> polyA\_site

<222> 484..495

<400> 244

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agtaggaasg cgccgscgt ggaggcgcca cgtcccttgc sgcggcggga gagamatcgc      60
ttggacttcg gggcggcctc ggaaggcc atg gcc ttt acc ctg tas tca ctg      112
                               Met Ala Phe Thr Leu Xaa Ser Leu
                               -10
ctg cag gca gcc ctg ctc tgc gtc aac gcc atc gca gtg ctg cac gag      160
Leu Gln Ala Ala Leu Leu Cys Val Asn Ala Ile Ala Val Leu His Glu
-5                               1                               5                               10
gag cga ttc ctc aag aac att ggc tgg gga aca gac cag gga att ggt      208
Glu Arg Phe Leu Lys Asn Ile Gly Trp Gly Thr Asp Gln Gly Ile Gly
                               15                               20                               25
gga ttt gga gaa gag ccg gga att aaa tca sag sta atg avs ctt att      256
Gly Phe Gly Glu Glu Pro Gly Ile Lys Ser Xaa Xaa Met Xaa Leu Ile
                               30                               35                               40
cga tct gta aga acc gtg atg aga gtg cca ttg ata ata gta aac tca      304
Arg Ser Val Arg Thr Val Met Arg Val Pro Leu Ile Ile Val Asn Ser
                               45                               50                               55
att gca att gtg tta ctt tta tta ttt gga tgaatwtcat tggagaaaat      354
Ile Ala Ile Val Leu Leu Leu Leu Phe Gly
60                               65
ggakactcag aaraggacat gccaktaraa kttattactt tggtcattat tggaatatatt      414
atatcttagc tggctgacct tgcacttgtc aaaaatgtaa agctgaaaat aaaaccaggg      474
tttctattta aaaaaaaaaa a      495

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<210> 245

<211> 884

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 21..614

<221> sig\_peptide

<222> 21..83

<223> Von Heijne matrix

score 10

seq LWALAMVTRPASA/AP

<221> polyA\_signal

<222> 849..854

<221> polyA\_site

<222> 873..884

<400> 245

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aataccttag accctcagtc atg cca gtg cct gct ctg tgc ctg ctc tgg gcc      53
                               Met Pro Val Pro Ala Leu Cys Leu Leu Trp Ala
                               -20                               -15
ctg gca atg gtg acc cgg cct gcc tca gcg gcc ccc atg ggc ggc cca      101
Leu Ala Met Val Thr Arg Pro Ala Ser Ala Ala Pro Met Gly Gly Pro
-10                               -5                               1                               5
gaa ctg gca cag cat gag gag ctg acc ctg ctc ttc cat ggg acc ctg      149

```

Glu	Leu	Ala	Gln	His	Glu	Glu	Leu	Thr	Leu	Leu	Phe	His	Gly	Thr	Leu		
			10					15					20				
cag	ctg	ggc	cag	gcc	ctc	aac	ggg	gtg	tac	agg	acc	acg	gag	gga	cgg	197	
Gln	Leu	Gly	Gln	Ala	Leu	Asn	Gly	Val	Tyr	Arg	Thr	Thr	Glu	Gly	Arg		
		25					30					35					
ctg	aca	aag	gcc	agg	aac	agc	ctg	ggg	ctc	tat	ggc	cgc	aca	ata	gaa	245	
Leu	Thr	Lys	Ala	Arg	Asn	Ser	Leu	Gly	Leu	Tyr	Gly	Arg	Thr	Ile	Glu		
		40				45					50						
ctc	ctg	ggg	cag	gag	gtc	agc	cgg	ggc	cgg	gat	gca	gcc	cag	gaa	ctt	293	
Leu	Leu	Gly	Gln	Glu	Val	Ser	Arg	Gly	Arg	Asp	Ala	Ala	Gln	Glu	Leu		
55					60					65				70			
cgg	gca	agc	ctg	ttg	gaa	act	car	atg	gag	gag	gat	att	ctg	cas	ctg	341	
Arg	Ala	Ser	Leu	Leu	Glu	Thr	Gln	Met	Glu	Glu	Asp	Ile	Leu	Xaa	Leu		
			75						80				85				
cag	gca	rag	gcc	aca	gct	gag	gtg	ctg	ggg	gag	gtg	gcc	cag	gca	car	389	
Gln	Ala	Xaa	Ala	Thr	Ala	Glu	Val	Leu	Gly	Glu	Val	Ala	Gln	Ala	Gln		
			90					95				100					
aag	gtg	cta	cgg	gac	agc	gtg	cag	cgg	cta	daa	ktc	cag	ctg	arg	asc	437	
Lys	Val	Leu	Arg	Asp	Ser	Val	Gln	Arg	Leu	Xaa	Xaa	Gln	Leu	Xaa	Xaa		
		105					110					115					
gcc	tgg	ctg	ggc	cct	gcc	tac	cga	aaa	ttt	gar	gtc	tta	aag	gcy	ccc	485	
Ala	Trp	Leu	Gly	Pro	Ala	Tyr	Arg	Lys	Phe	Glu	Val	Leu	Lys	Ala	Pro		
		120				125					130						
ck	gam	aar	car	aac	cac	atc	cta	tgg	gcc	ctc	aca	ggc	cac	gtg	cak	533	
Pro	Xaa	Lys	Gln	Asn	His	Ile	Leu	Trp	Ala	Leu	Thr	Gly	His	Val	Xaa		
		135				140					145			150			
cgg	car	arg	cgg	gar	atg	gtg	gca	cag	cag	cwt	ckg	ctg	cna	car	atc	581	
Arg	Gln	Xaa	Arg	Glu	Met	Val	Ala	Gln	Gln	Xaa	Xaa	Leu	Xaa	Gln	Ile		
			155					160				165					
cag	gar	aaa	ctc	cac	aca	gcg	gcg	ctc	cca	gcc	tgaatctgcc	tggaatggaac				634	
Gln	Glu	Lys	Leu	His	Thr	Ala	Ala	Leu	Pro	Ala							
		170						175									
tgaggaccaa	tcattgctgca	aggaacactt	ccacgccccg	tgaggccccct	gtgcagggag											694	
gagctgcttg	ttcactggga	tcagccaggg	cgccggggcc	cacttctgag	cacagagcar											754	
agacagacgc	aggcggggac	aaaggcagag	gatgtagccc	cattggggag	gggtggagga											814	
aggacatgta	ccctttcatr	mctacacacc	cctcattaaa	gcavagtcgt	ggcatctcaa											874	
aaaaaaaaaa																884	

<210> 246  
 <211> 897  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 94..573

<221> sig\_peptide  
 <222> 94..258  
 <223> Von Heijne matrix  
 score 4.69999980926514  
 seq IGILCSLLGTVLL/WV

<221> polyA\_signal  
 <222> 862..867

<221> polyA\_site

<222> 886..897

<400> 246

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aaggcgcgct gcctagcacc cggaagagcc gtcaacttag cgagcgcaac aggctgccgc      60
tgaggagctg gagctggtgg ggactgggccc gca atg gac aag ctg aag aag gtg      114
                               Met Asp Lys Leu Lys Lys Val
                               -55                               -50

ctg agc ggg cag gac acg gag gac cgg agc ggc ctg tcc gag gtt gtt      162
Leu Ser Gly Gln Asp Thr Glu Asp Arg Ser Gly Leu Ser Glu Val Val
                               -45                               -40                               -35

gag gca tct tca tta agc tgg agt acc agg ata aaa ggc ttc att gcg      210
Glu Ala Ser Ser Leu Ser Trp Ser Thr Arg Ile Lys Gly Phe Ile Ala
                               -30                               -25                               -20

tgt ttt gct ata gga att ctc tgc tca ctg ctg ggt act gtt ctg ctg      258
Cys Phe Ala Ile Gly Ile Leu Cys Ser Leu Leu Gly Thr Val Leu Leu
                               -15                               -10                               -5

tgg gtg ccc agg aag gga cta cac ctc ttc gca gtg ttt tat acc ttt      306
Trp Val Pro Arg Lys Gly Leu His Leu Phe Ala Val Phe Tyr Thr Phe
1                               5                               10                               15

ggg aat atc gca tca att ggg agt acc atc ttc ctc atg gga cca gtg      354
Gly Asn Ile Ala Ser Ile Gly Ser Thr Ile Phe Leu Met Gly Pro Val
                               20                               25                               30

aaa cag ctg aag cga atg ttt gag cct act cgt ttg att gca act atc      402
Lys Gln Leu Lys Arg Met Phe Glu Pro Thr Arg Leu Ile Ala Thr Ile
                               35                               40                               45

atg gtg ctg ttg tgt ttt gca ctt acc ctg tgt tct gcc ttt tgg tgg      450
Met Val Leu Leu Cys Phe Ala Leu Thr Leu Cys Ser Ala Phe Trp Trp
50                               55                               60

cat aac aag gga ctt gca ctt atc ttc tgc att ttg cag tct ttg gca      498
His Asn Lys Gly Leu Ala Leu Ile Phe Cys Ile Leu Gln Ser Leu Ala
65                               70                               75                               80

ttg acg tgg tac agc ctt tcc ttc ata cca ttt gca agg gat gct gtg      546
Leu Thr Trp Tyr Ser Leu Ser Phe Ile Pro Phe Ala Arg Asp Ala Val
                               85                               90                               95

aaa aad tgt ttt gcc gtg tgt ctt gca taattcatgg ccagttttat      593
Lys Xaa Cys Phe Ala Val Cys Leu Ala
100                               105

gaagcttttg aaggcactat ggacagaagc tgggtggacag ttttgtwact atcttcgaaa      653
cctctgtctt acagacatgt gccttttatc ttgcagcaat gtgttgcttg tgattcgaac      713
atttgagggg tactttttgga agcaacaata cattctcgaa cctgaatgtc agtagcacag      773
gatgagaagt gggttctgta tcttgtggag tggaatcttc ctcatgtacc tgtttcctct      833
ctggatgttg tccactgaa ttcccatgaa tacaaaccta ttcagcaaca gcaaaaaaaaa      893
aaaa                                                                897

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<210> 247

<211> 518

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 74..397



<221> sig\_peptide  
<222> 74..127  
<223> Von Heijne matrix  
score 7.69999980926514  
seq LLLLPVLGLLVSS/KT

<221> polyA\_signal  
<222> 472..477

<221> polyA\_site  
<222> 507..518

<400> 247  
aaagaaagag ctgcsgtgca ggaattcgtg tgccggattt ggtagctga gccaccgag 60  
aggcgctctg agg atg aaa gct ctc tgt ctc ctc ctc ctc cct gtc ctg 109  
Met Lys Ala Leu Cys Leu Leu Leu Leu Pro Val Leu  
-15 -10  
ggg ctg ttg gtg tct agc aag acc ctg tgc tcc atg gaa gaa gcc atc 157  
Gly Leu Leu Val Ser Ser Lys Thr Leu Cys Ser Met Glu Glu Ala Ile  
-5 1 5 10  
aat gag agg atc cag gag gtc gcc ggc tcc cta ata ttt agg gca ata 205  
Asn Glu Arg Ile Gln Glu Val Ala Gly Ser Leu Ile Phe Arg Ala Ile  
15 20 25  
agc agc att ggc cga ggg agc gag agc gtc acc tcc agg ggg gac ctg 253  
Ser Ser Ile Gly Arg Gly Ser Glu Ser Val Thr Ser Arg Gly Asp Leu  
30 35 40  
gct act tgc ccc cga ggc ttc gcc gtc acc ggc tgc act tgt ggc tcc 301  
Ala Thr Cys Pro Arg Gly Phe Ala Val Thr Gly Cys Thr Cys Gly Ser  
45 50 55  
gcc tgt ggc tcg tgg gat gtg cgc gcc gag acc aca tgt cac tgc cag 349  
Ala Cys Gly Ser Trp Asp Val Arg Ala Glu Thr Thr Cys His Cys Gln  
60 65 70  
tgc gcg ggc atg gac tgg acc gga gcg cgc tgc tgt cgt gtg cag ccc 397  
Cys Ala Gly Met Asp Trp Thr Gly Ala Arg Cys Cys Arg Val Gln Pro  
75 80 85 90  
tgaggtcgcg cgcagcgcgt gcacagcgcg ggcggaggcg gctccagggtc cggagggggtt 457  
gcggggggagc tggaaataaa cctggagatg atgatgatga tgatgatgga aaaaaaaaaa 517  
a 518

<210> 248  
<211> 350  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 51..242

<221> sig\_peptide  
<222> 51..116  
<223> Von Heijne matrix  
score 6.5  
seq SCLCPALFPGTSS/FI

<221> polyA\_signal

<222> 319..324

<221> polyA\_site

<222> 339..350

<400> 248

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acgtcattcc aaaaccacac ccttgcaaag ctttgtactc cgcaccccag atg atc      56
                                     Met Ile
tcc agg cag ctc aga tct ctt tcc tgc ctt tgc cct gca ctg ttc ccc      104
Ser Arg Gln Leu Arg Ser Leu Ser Cys Leu Cys Pro Ala Leu Phe Pro
-20          -15          -10          -5
ggg act tcc tcc ttt att gta gca ctc agc tcc cca gcc gat ctg tac      152
Gly Thr Ser Ser Phe Ile Val Ala Leu Ser Ser Pro Ala Asp Leu Tyr
          1          5          10
atc cct cav agg cas cga tct gat gaa ttg gtt ttt gaa tcc car aaa      200
Ile Pro Xaa Arg Xaa Arg Ser Asp Glu Leu Val Phe Glu Ser Gln Lys
          15          20          25
ggg tct gcc atg gag ttg gca gtc atc acg gta rat ggc gta      242
Gly Ser Ala Met Glu Leu Ala Val Ile Thr Val Xaa Gly Val
          30          35          40
tgattttgct gaattttaaa taaaatgaaa accataaatt acatratgct tttattgach      302
cttgacmact ggcctaaata aaaaractct gactccaaaa aaaaaaaaa      350
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<210> 249

<211> 996

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 111..191

<221> sig\_peptide

<222> 111..155

<223> Von Heijne matrix

score 5.80000019073486

seq FLXLMTLTTHVHS/SA

<221> polyA\_signal

<222> 965..970

<221> polyA\_site

<222> 986..996

<400> 249

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atccgataca gaacatgcag taatgtggac tgcccaccag aagcaggtga tttccgagct      60
cagcaatgct cagctcataa tgatgtcaag caccatggcc agttttatga atg ggy      116
                                     Met Gly
                                     -15
ttc ctg wgt cta atg acc ctg aca acc cat gtt cac tca agt gcc aag      164
Phe Leu Xaa Leu Met Thr Leu Thr Thr His Val His Ser Ser Ala Lys
          -10          -5          1
cca aat gaa caa ccc tgg ttg ttg aac tagcacctaa ggtcttarat      211
Pro Asn Glu Gln Pro Trp Leu Leu Asn
          5          10
```

```
ggtagcggtt gctatacaga atctttggat atgtgcatca gtggtttatg ccaaattggt 271
ggctgcgatac accagctggg aagcaccgtc aaggaarata actgtggggg ctgcaacrga 331
natgggtcca cctgccggct ggtccgaggg cartataaat cccakctctc cgcaacccaaa 391
tcrgatgata ctgtggttgc aattccctat ggaagtakac atattcgctt tgtcttaaaa 451
ggtcctgatac acttatatct ggaareccawa accctccagg ggactaawgg tgaaaacagt 511
ctcasctcca caggaacttt ccttgtggac aattctagt tggacttcca gaawtttcca 571
gacwdagaga tactgagaat ggctggacca ctcacagcag atttcattgt caawattcgt 631
aactcgggct ccgctgacag tacagtccag kkcattctct atcaacccat catccaccga 691
tggagggara cggatttctt tccttgtctc gcaacctgtg gaggagggtta tcagctgaca 751
tcggctgagt gctacgatct gaggagcaac cgtgtgggtg ctgaccaata ctgtcactat 811
taccagaga acatcaaacc caaacccaag cttcaggagt gcaacttggg tccttgtcca 871
gccagggtcag tcaaatttgc tagttcattt gtcataaaca taactcaagt tccaaatagg 931
ttatttaaata taaaatgaaa cgttttaatt aaaaataaaa tgaaattaaa catcaaaaaa 991
aaaaa 996
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<210> 250  
<211> 860  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 45..602  
<221> sig\_peptide  
<222> 45..107  
<223> Von Heijne matrix  
score 8.5  
seq LLTIVGLILPTRG/QT  
<221> polyA\_signal  
<222> 828..833  
<221> polyA\_site  
<222> 850..860

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<400> 250
acctctctcc acgaggctgc cggcttagga cccccagctc cgac atg tcg ccc tct 56
Met Ser Pro Ser
-20
ggt cgc ctg tgt ctt ctc acc atc gtt ggc ctg att ctc ccc acc aga 104
Gly Arg Leu Cys Leu Leu Thr Ile Val Gly Leu Ile Leu Pro Thr Arg
-15 -10 -5
gga cag acg ttg aaa gat acc acg tcc agt tct tca gca gac tca act 152
Gly Gln Thr Leu Lys Asp Thr Thr Ser Ser Ser Ser Ala Asp Ser Thr
1 5 10 15
atc atg gac att cag gtc ccg aca cga gcc cca gat gca gtc tac aca 200
Ile Met Asp Ile Gln Val Pro Thr Arg Ala Pro Asp Ala Val Tyr Thr
20 25 30
gaa ctc cag ccc acc tct cca acc cca acc tgg cct gct gat gaa aca 248
Glu Leu Gln Pro Thr Ser Pro Thr Pro Thr Trp Pro Ala Asp Glu Thr
35 40 45
cca caa ccc cag acc cag acc cag caa ctg gaa gga acg gat ggg cct 296
Pro Gln Pro Gln Thr Gln Thr Gln Gln Leu Glu Gly Thr Asp Gly Pro
50 55 60
```

cta gtg aca gat cca gag aca cac wak agc mcc aaa gca gct cat ccc	344
Leu Val Thr Asp Pro Glu Thr His Xaa Ser Xaa Lys Ala Ala His Pro	
65 70 75	
act gat gac acc acg acg ctc tct gag aga cca tcc cca agc aca kac	392
Thr Asp Asp Thr Thr Thr Leu Ser Glu Arg Pro Ser Pro Ser Thr Xaa	
80 85 90 95	
gtc cat dac aga ccb cba kda ccc tca akc cat ctg gtt ttc atg agg	440
Val His Xaa Arg Pro Xaa Xaa Pro Ser Xaa His Leu Val Phe Met Arg	
100 105 110	
atg acc cct tct tct atg atg aac aca ccc tcc gga aac sgg ggc tgt	488
Met Thr Pro Ser Ser Met Met Asn Thr Pro Ser Gly Asn Xaa Gly Cys	
115 120 125	
tgg tcg cag ctg tgc tgt tca tca cag gca tca tca tcc tca cca gtg	536
Trp Ser Gln Leu Cys Cys Ser Ser Gln Ala Ser Ser Ser Ser Pro Val	
130 135 140	
gca agt gca ggc agc tgt ccc ggt tat gcc gga atc att gca ggt gag	584
Ala Ser Ala Gly Ser Cys Pro Gly Tyr Ala Gly Ile Ile Ala Gly Glu	
145 150 155	
tcc atc aga aac agg agc tgacaacctg ctgggcaccc gaagaccaag	632
Ser Ile Arg Asn Arg Ser	
160 165	
ccccctgccca gctcaccgtg cccagcctcc tgcacccct cgaagagcct ggccagagag	692
ggaagacaca gatgatgaag ctggagccag ggctgccggt ccgagtcctcc tacctcccc	752
aaccctgccc gcccctgaag gctacctggc gccttggggg ctgtccctca agttatctcc	812
cttgctaaga caaaaagtaa agcactgtgg tctttgcaaa aaaaaaaaa	860

<210> 251

<211> 593

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 24..560

<221> sig\_peptide

<222> 24..101

<223> Von Heijne matrix

score 10.3999996185303

seq LLLLLCGPSQDQC/RP

<221> polyA\_signal

<222> 563..568

<221> polyA\_site

<222> 583..593

<400> 251

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ctt ctc ccg ttg ctg ctg ctg ctc tgc ggc cct tcc cag gat caa tgc	101
Leu Leu Pro Leu Leu Leu Leu Leu Cys Gly Pro Ser Gln Asp Gln Cys	
-15 -10 -5	

cga cct gta ctc cag aat ctg ttg cag agc cca ggc ttg aca tgg agc	149
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Arg Pro Val Leu Gln Asn Leu Leu Gln Ser Pro Gly Leu Thr Trp Ser
1          5          10          15
ttg gaa gtg ccc act ggg aga gaa gga aag gaa ggt ggg gat cgg gga    197
Leu Glu Val Pro Thr Gly Arg Glu Gly Lys Glu Gly Gly Asp Arg Gly
          20          25          30
cca ggg cta akt ggg gcc act cca gcc agg agc cct cag ggc aag gag    245
Pro Gly Leu Xaa Gly Ala Thr Pro Ala Arg Ser Pro Gln Gly Lys Glu
          35          40          45
atg ggg aga caa agg acc aga aag gtg aag ggc cct gct tgg akt cac    293
Met Gly Arg Gln Arg Thr Arg Lys Val Lys Gly Pro Ala Trp Xaa His
          50          55          60
aca gca aat cag gaa cta aac agg atg agg tct ctg tct tct ggc tcc    341
Thr Ala Asn Gln Glu Leu Asn Arg Met Arg Ser Leu Ser Ser Gly Ser
          65          70          75          80
gtg cca gtg ggg cat ctg gag ggt ggc acg gtc aag ctt cag aag gac    389
Val Pro Val Gly His Leu Glu Gly Gly Thr Val Lys Leu Gln Lys Asp
          85          90          95
acg ggc ctc cat tcc tgc ara gat ggt atg gct tct ctt gaa ggg acg    437
Thr Gly Leu His Ser Cys Xaa Asp Gly Met Ala Ser Leu Glu Gly Thr
          100          105          110
cca gct tca gtc ctg gct gat gct tgc cca gga ttc cat gat gtg aan    485
Pro Ala Ser Val Leu Ala Asp Ala Cys Pro Gly Phe His Asp Val Xaa
          115          120          125
gtt car arg gcc cta ttt ggg tta agt ggg ana rta ctg tgg ctg aaa    533
Val Gln Xaa Ala Leu Phe Gly Leu Ser Gly Xaa Xaa Leu Trp Leu Lys
          130          135          140
acc cac ttc tgc ctt tct att ana ctt taaataaact ctgaaracct    580
Thr His Phe Cys Leu Ser Ile Xaa Leu
          145          150
gtataaaaaaaaaaaa aaa    593

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      seq VAFMLTLPILVCK/VQ

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<222> 1104..1114

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                        Met Ala Thr
                        -55
aca gtc cct gat ggt tgc cgc aat ggc ctg aaa tcc aag tac tac aga    165

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Thr	Val	Pro	Asp	Gly	Cys	Arg	Asn	Gly	Leu	Lys	Ser	Lys	Tyr	Tyr	Arg		
		-50					-45					-40					
ctt	tgt	gat	aag	gct	gaa	gct	tgg	ggc	atc	gtc	cta	gaa	acg	gtg	gcc	213	
Leu	Cys	Asp	Lys	Ala	Glu	Ala	Trp	Gly	Ile	Val	Leu	Glu	Thr	Val	Ala		
		-35				-30					-25						
aca	gcc	ggg	gtt	gtg	acc	tgc	gtg	gcc	ttc	atg	ctg	act	ctc	ccg	atc	261	
Thr	Ala	Gly	Val	Val	Thr	Ser	Val	Ala	Phe	Met	Leu	Thr	Leu	Pro	Ile		
		-20			-15				-10					-5			
ctc	gtc	tgc	aag	gtg	cag	gac	tcc	aac	agg	cga	aaa	atg	ctg	cct	act	309	
Leu	Val	Cys	Lys	Val	Gln	Asp	Ser	Asn	Arg	Arg	Lys	Met	Leu	Pro	Thr		
			1				5				10						
cag	ttt	ctc	ttc	ctc	ctg	ggg	gtg	ttg	ggc	atc	ttt	ggc	ctc	acc	ttc	357	
Gln	Phe	Leu	Phe	Leu	Leu	Gly	Val	Leu	Gly	Ile	Phe	Gly	Leu	Thr	Phe		
		15				20					25						
gcc	ttc	atc	atc	gga	ctg	gac	ggg	agc	aca	ggg	ccc	aca	cgc	ttc	ttc	405	
Ala	Phe	Ile	Ile	Gly	Leu	Asp	Gly	Ser	Thr	Gly	Pro	Thr	Arg	Phe	Phe		
		30				35					40						
ctc	ttt	ggg	atc	ctc	ttt	tcc	atc	tgc	ttc	tcc	tgc	ctg	ctg	gct	cat	453	
Leu	Phe	Gly	Ile	Leu	Phe	Ser	Ile	Cys	Phe	Ser	Cys	Leu	Leu	Ala	His		
		45			50				55					60			
gct	gtc	agt	ctg	acc	aag	ctc	gtc	cgg	ggg	agg	aaa	gcc	cct	ttc	cct	501	
Ala	Val	Ser	Leu	Thr	Lys	Leu	Val	Arg	Gly	Arg	Lys	Ala	Pro	Phe	Pro		
			65				70						75				
gtt	ggt	gat	tct	ggg	tct	ggc	cgt	ggg	ctt	cag	cct	agt	cca	gga	tgt	549	
Val	Gly	Asp	Ser	Gly	Ser	Gly	Arg	Gly	Leu	Gln	Pro	Ser	Pro	Gly	Cys		
			80				85				90						
tat	cgc	tat	tgaatatatt	gtcctgacca	tgaataggac	caacgtcaat										598	
Tyr	Arg	Tyr															
		95															
gtctttttctg	agctttccgc	tcctcgctgc	aatgaaaact	ttgtcctcct	gctcacctac											658	
ktcctctttct	tgatggcgct	gaccttctc	wtgtcctcct	tcaccttctg	tggtkccttc											718	
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gagttttggc	tgctcacaac	gcaackaaac	cccatggatt	atcctgttga	ggatgctttc											958	
tgtaaacctc	aactcgtgaa	gaagagctat	ggtgtggrga	acagagccta	skctcaagag											1018	
gaaatcactc	aaggttttga	agagacagg	gacacgctct	atgccccta	ttccacacat											1078	
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<211> 1182

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<222> 128..220

<223> Von Heijne matrix

score 4.69999980926514

seq LAVDSWWLDPGHA/AV

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<222> 1145..1150

<221> polyA\_site

<222> 1170..1181

<400> 253

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ccgcgcgcgt	ttcctg	atttg	gtt	gtg	gggtg	gctac	ctctt	cg	ttctg	att	ggccg	ctagt	120			
gagcaag	atg	ctg	agc	aag	ggt	ctg	aag	cg	aaa	cg	gag	gag	gag	169		
	Met	Leu	Ser	Lys	Gly	Leu	Lys	Arg	Lys	Arg	Glu	Glu	Glu			
		-30				-25				-20						
gag	aag	gaa	cct	ctg	gca	gtc	gac	tcc	tgg	tgg	cta	gat	cct	ggc	cac	217
Glu	Lys	Glu	Pro	Leu	Ala	Val	Asp	Ser	Trp	Trp	Leu	Asp	Pro	Gly	His	
		-15				-10				-5						
gca	gcg	gtg	gca	cag	gca	ccc	ccg	gcc	gtg	gcc	tct	agc	tcc	ctc	ttt	265
Ala	Ala	Val	Ala	Gln	Ala	Pro	Pro	Ala	Val	Ala	Ser	Ser	Ser	Leu	Phe	
	1			5					10					15		
gac	ctc	tca	gtg	ctc	aag	ctc	cac	cac	agc	ctg	cag	vrr	agt	rag	ccg	313
Asp	Leu	Ser	Val	Leu	Lys	Leu	His	His	Ser	Leu	Gln	Xaa	Ser	Xaa	Pro	
			20						25				30			
gac	ctg	cg	cac	ctg	gtg	ctg	gtc	atr	aac	act	ctg	cg	cg	atc	cag	361
Asp	Leu	Arg	His	Leu	Val	Leu	Val	Xaa	Asn	Thr	Leu	Arg	Arg	Ile	Gln	
			35					40					45			
gcg	tcc	atg	gca	ccc	gcg	gct	gcc	ctg	cca	cct	gtg	cct	acc	cca	cct	409
Ala	Ser	Met	Ala	Pro	Ala	Ala	Ala	Leu	Pro	Pro	Val	Pro	Thr	Pro	Pro	
		50				55					60					
gca	gcc	ccc	ant	gtg	gct	gac	aac	tta	ctg	gca	agc	tcg	gac	gct	gcc	457
Ala	Ala	Pro	Xaa	Val	Ala	Asp	Asn	Leu	Leu	Ala	Ser	Ser	Asp	Ala	Ala	
		65				70				75						
ctt	tca	gcc	tcc	atg	gcc	arm	ctc	ctg	gar	gac	ctc	agc	cac	att	gag	505
Leu	Ser	Ala	Ser	Met	Ala	Xaa	Leu	Leu	Glu	Asp	Leu	Ser	His	Ile	Glu	
		80				85				90				95		
ggc	ctg	agt	cag	gct	ccc	caa	ccc	ttg	gca	gac	gag	ggg	cca	cca	ggc	553
Gly	Leu	Ser	Gln	Ala	Pro	Gln	Pro	Leu	Ala	Asp	Glu	Gly	Pro	Pro	Gly	
			100						105				110			
cg	agc	atc	ggg	gga	wca	ccg	ccc	amc	ctg	gg	gcc	ttg	gac	ctg	ctg	601
Arg	Ser	Ile	Gly	Gly	Xaa	Pro	Pro	Xaa	Leu	Gly	Ala	Leu	Asp	Leu	Leu	
			115					120					125			
ggc	cca	gcc	act	ggc	tgt	cta	ctg	gac	aat	ggg	ctt	gag	ggc	ctg	ttt	649
Gly	Pro	Ala	Thr	Gly	Cys	Leu	Leu	Asp	Asn	Gly	Leu	Glu	Gly	Leu	Phe	
		130				135						140				
gag	gat	att	gac	acc	tct	atg	tat	gac	aat	gaa	ctt	tgg	gca	cca	gcc	697
Glu	Asp	Ile	Asp	Thr	Ser	Met	Tyr	Asp	Asn	Glu	Leu	Trp	Ala	Pro	Ala	
		145				150					155					
tct	gag	ggc	ctc	aaa	cca	ggc	cct	gag	gat	ggg	ccg	ggc	aag	gag	gaa	745
Ser	Glu	Gly	Leu	Lys	Pro	Gly	Pro	Glu	Asp	Gly	Pro	Gly	Lys	Glu	Glu	
		160				165				170				175		
gct	ccg	gag	ctg	gac	gag	gcc	gaa	ttg	gac	tac	ctc	atg	gat	gtg	ctg	793
Ala	Pro	Glu	Leu	Asp	Glu	Ala	Glu	Leu	Asp	Tyr	Leu	Met	Asp	Val	Leu	
			180						185					190		
gtg	ggc	aca	cag	gca	ctg	gag	cga	ccg	ccg	ggg	cca	ggg	cg			835
Val	Gly	Thr	Gln	Ala	Leu	Glu	Arg	Pro	Pro	Gly	Pro	Gly	Arg			
			195					200				205				
tgagcc	ctcg	tgctg	gaatg	gtt	gtct	gggt	atct	gaactg	agc	ctg	ctgg	accaac				895
tg	ctctcg	aa	gacac	agc	tg	gcttc	ccct	ag	tacag	aga	acag	ggctg	ggcc	actttg		955
gag	agac	aga	atct	agtc	ctt	ggg	caact	tc	acat	ccgt	cc	tctgt	ctca	ggg	ctggc	1015
ggg	gag	cc	ctg	gaatt	acccc	ctag	t	gatgg	aat	gac	agg	tct	gtgg	ggg	act	1075

ctggccctgg ggtcatagct tgggctgttc cttctctgat acggaagag acccaatcag 1135  
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 score 3.70000004768372  
 seq LASSFLFTMGGLG/FI

<221> polyA\_signal  
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 <222> 1062..1073

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 atg gag act ttg tac cgt gtc ccg ttc tta gtg ctc gaa tgt ccc aac 106  
 Met Glu Thr Leu Tyr Arg Val Pro Phe Leu Val Leu Glu Cys Pro Asn  
 -100 -95 -90 -85  
 ctg aag ctg aag aag ccg ccc tgg ttg cac atg ccg tcg gcc atg act 154  
 Leu Lys Leu Lys Lys Pro Pro Trp Leu His Met Pro Ser Ala Met Thr  
 -80 -75 -70  
 gtg tat gct ctg gtg gtg gtg tct tac ttc ctc atc acc gga gga ata 202  
 Val Tyr Ala Leu Val Val Val Ser Tyr Phe Leu Ile Thr Gly Gly Ile  
 -65 -60 -55  
 att tat gat gtt att gtt gaa cct cca agt gtc ggt tct atg act gat 250  
 Ile Tyr Asp Val Ile Val Glu Pro Pro Ser Val Gly Ser Met Thr Asp  
 -50 -45 -40  
 gaa cat ggg cat cag agg cca gta gct ttc ttg gcc tac aga gta aat 298  
 Glu His Gly His Gln Arg Pro Val Ala Phe Leu Ala Tyr Arg Val Asn  
 -35 -30 -25  
 gga caa tat att atg gaa gga ctt gca tcc agc ttc cta ttt aca atg 346  
 Gly Gln Tyr Ile Met Glu Gly Leu Ala Ser Ser Phe Leu Phe Thr Met  
 -20 -15 -10 -5  
 gga ggt tta ggt ttc ata atc ctg gac gga tcg aat gca cca aat atc 394  
 Gly Gly Leu Gly Phe Ile Ile Leu Asp Gly Ser Asn Ala Pro Asn Ile  
 1 5 10  
 cca aaa ctc aat aga ttc ctt ctt ctg ttc att gga ttc gtc tgt gtc 442  
 Pro Lys Leu Asn Arg Phe Leu Leu Phe Ile Gly Phe Val Cys Val  
 15 20 25  
 cta twr agt ttt ttc ayy gct aga gta ttc atg aga atg aaa ctg ccg 490  
 Leu Xaa Ser Phe Xaa Xaa Ala Arg Val Phe Met Arg Met Lys Leu Pro  
 30 35 40  
 ggc tat ctg atg ggt tagagtgcct ttgasaagaa atcagtggat actggatttg 545  
 Gly Tyr Leu Met Gly



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45
ctcctgtcaa wgaastttta aaggctgtmc caatcctcta atatgaaatg tggaaaagaa 605
tgaagagcag cagtaaaaga aatatctagt gaaaaaacag gaagcgtatt gaagcttgga 665
ctagaatttc ttcttgggat taaagagaca agtttatcac agaatttttt ttctgtctgg 725
cctattgcta taccaatgat gttgagtggc attttctttt tagtttttca ttaaaatata 785
ttccatatct acaactataa tatcaaataa agtgattatt ttttacaacc ctcttaacat 845
tttttgagaga tgacatttct gattttcaga aattaacata aaatccagaa gcaagattcc 905
gtaagctgag aactctggac agttgatcag ctttacctat ggtgctttgc ctttaactag 965
agtggtgat ggtagattat ttcagatatg tatgtaaaac tgtttcctga acaataagat 1025
gtatgaacgg agcagaaata aatacttttt ctaattaaaa aaaaaaaaa 1073

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<210> 255
<211> 818
<212> DNA
<213> Homo sapiens

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<222> 1..207

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<221> sig_peptide
<222> 1..147
<223> Von Heijne matrix
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      seq HLPFLLLLSCVGX/XP

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<221> polyA_signal
<222> 784..789

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<221> polyA_site
<222> 807..818

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Met Pro Phe His Phe Pro Phe Leu Gly Phe Val Cys Leu His Leu His
      -45      -40      -35
ctt acc cct tgc ctg act gta ccc cgt aga ccc ctg ttt ctc ctc ctg 96
Leu Thr Pro Cys Leu Thr Val Pro Arg Arg Pro Leu Phe Leu Leu Leu
      -30      -25      -20
cac ctg tgt ccc cat ctg ccc ttc ttg ttg ctc ctg tca tgt gtc ggg 144
His Leu Cys Pro His Leu Pro Phe Leu Leu Leu Leu Ser Cys Val Gly
      -15      -10      -5
gkc www ccc tcc tgt ctg cct tct tcc tcc act tgt gtc agc ttg cat 192
Xaa Xaa Pro Ser Cys Leu Pro Ser Ser Ser Thr Cys Val Ser Leu His
      1      5      10      15
ttt ttt att cct gac tgagtcacca caccctctc cctgatcaa agggaatatk 247
Phe Phe Ile Pro Asp
      20
artttttaat ttggatcgac tgaggtgccca ggagaaactg cagkcccagg tatccmvaca 307
gccaccagga tggtcctctg cccaccccc accgcctctk cccaccttt tccaacgtgt 367
tgcattgctgg gaactggggg gtgtggggga aggggctgcc ggcttctttc aggangctga 427
rgtttggar gaaaatcaac ctgggaracc acccggccg cggcgctca gtggacaggt 487
gggargaaaa gaaaacttct taccttggar garggacatc ccgcttcctt atccttagct 547
tttttgttgc tctctccac tgccctttt aatttatttg gttgtttgcg gaaggagggg 607
ggaagggggg aagctggggc gggaactgtc cgaggtgctg agctggggcg ggaccggaat 667

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cctccccgcta ggggtaccagg gactgagttg ggccctggggc cgtgtccaag gtgccaatga 727
tgcggggccga cagarcgggc cgcactgtct gtctgtccgt ctgtcccga aagaactata 787
aagcgctgga agcgctgca aaaaaaaaaa a 818
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<212> DNA  
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<222> 12..734

<221> sig\_peptide  
<222> 12..101  
<223> Von Heijne matrix  
score 4.80000019073486  
seq ILFCVGAVGACTL/SV

<221> polyA\_signal  
<222> 914..919  
<221> polyA\_site  
<222> 961..971

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Met Gly Thr Ala Ser Arg Ser Asn Ile Ala Arg His Leu
-30 -25 -20
caa acc aat ctc att cta ttt tgt gtc ggt gct gtg ggc gcc tgt act 98
Gln Thr Asn Leu Ile Leu Phe Cys Val Gly Ala Val Gly Ala Cys Thr
-15 -10 -5
ctc tct gtc aca caa ccg tgg tac cta gaa gtg gac tac act cat gag 146
Leu Ser Val Thr Gln Pro Trp Tyr Leu Glu Val Asp Tyr Thr His Glu
1 5 10 15
gcc gtc acc ata aag tgt acc ttc tcc gca acc gga tgc cct tct gag 194
Ala Val Thr Ile Lys Cys Thr Phe Ser Ala Thr Gly Cys Pro Ser Glu
20 25 30
caa cca aca tgc ctg tgg ttt cgc tac ggt gct cac cag cct gag aac 242
Gln Pro Thr Cys Leu Trp Phe Arg Tyr Gly Ala His Gln Pro Glu Asn
35 40 45
ctg tgc ttg gac ggg tgc aaa agt gag gca gas aag ttc aca gtg agg 290
Leu Cys Leu Asp Gly Cys Lys Ser Glu Ala Xaa Lys Phe Thr Val Arg
50 55 60
gag gcc ctc aaa gaa aac caa gtt tcc ctc act gta aac aga gtg act 338
Glu Ala Leu Lys Glu Asn Gln Val Ser Leu Thr Val Asn Arg Val Thr
65 70 75
tca aat gac agt gca att tac atc tgt gga ata gca ttc ccc agt gtg 386
Ser Asn Asp Ser Ala Ile Tyr Ile Cys Gly Ile Ala Phe Pro Ser Val
80 85 90 95
ccg gaa gcg aga gct aaa cag aca gga gga ggg acc aca ctg gtg gta 434
Pro Glu Ala Arg Ala Lys Gln Thr Gly Gly Gly Thr Thr Leu Val Val
100 105 110
aga gaa att aag ctg ctc agc aag gaa ctg cgg agc ttc ctg aca gct 482
Arg Glu Ile Lys Leu Leu Ser Lys Glu Leu Arg Ser Phe Leu Thr Ala
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      115              120              125
ctt gta tca ctg ctc tct gtc tat gtg acc ggt gtg tgc gtg gcc ttc      530
Leu Val Ser Leu Leu Ser Val Tyr Val Thr Gly Val Cys Val Ala Phe
      130              135              140
ata ctc ctc tcc aaa tca aaa tcc aac cct cta aga aac aaa gaa ata      578
Ile Leu Leu Ser Lys Ser Lys Ser Asn Pro Leu Arg Asn Lys Glu Ile
      145              150              155
aaa gaa gac tca caa aag aag aag agt gct cgg cgt att ttt cag gaa      626
Lys Glu Asp Ser Gln Lys Lys Lys Ser Ala Arg Arg Ile Phe Gln Glu
      160              165              170              175
att gct caa gaa cta tac cat aag aga cat gtg gaa aca aat cag caa      674
Ile Ala Gln Glu Leu Tyr His Lys Arg His Val Glu Thr Asn Gln Gln
      180              185              190
tct gag aaa gat aac aac act tat gaa aac aga aga gta ctt tcc aac      722
Ser Glu Lys Asp Asn Asn Thr Tyr Glu Asn Arg Arg Val Leu Ser Asn
      195              200              205
tat gaa agg cca tagaaacggt ttaattttca atgaagtcac tgaaaatcca      774
Tyr Glu Arg Pro
      210
actccaggag ctatggcagt gttaatgaac atatatcatc aggtcttaaa aaaaaataaa      834
ggtaaaactga aaagacaact ggctacaaag aaggatgcc aatgtgaagg aaactataac      894
taataktcat taccaaaata ctaaaaccca acaaaatgca actgaaaaat accttccaaa      954
tttgccaaaa aaaaaaw      971

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<210> 257

<211> 640

<212> DNA

<213> Homo sapiens

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<221> CDS

<222> 378..518

<221> sig\_peptide

<222> 378..467

<223> Von Heijne matrix

score 5.5

seq SLMTCTTLINASA/IS

<221> polyA\_signal

<222> 607..612

<221> polyA\_site

<222> 628..640

<400> 257

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aatggcaggg aacctcttat cgggggcagg taggcgctg tgggactggg tgccctctggc      180
gtgcagaagc ttctctcttg gtgtgcctag attgatcggg ataaggctca ctctcccgcc      240
cccaaagtg gttgatcgtt ggaacgagaa aagggccatg ttcggagtgt atgacaacat      300
cgggatcctg ggaaactttg aaaagcacc caaagaactg atcagggggc ccatatggct      360
tcgaggttgg aaaggga atg aat tgc aac gtt gta tcc gaa aga gga aaa      410

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Met Asn Cys Asn Val Val Ser Glu Arg Gly Lys

-30

-25

-20

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tgg ttg gaa gta gaa tgt tgc ctg atg acc tgc aca acc tta ata aac      458
Trp Leu Glu Val Glu Cys Ser Leu Met Thr Cys Thr Thr Leu Ile Asn
          -15                      -10                      -5
gca tcc gct atc tct aca aac act tta acc gac atg gga agt ttc gat      506
Ala Ser Ala Ile Ser Thr Asn Thr Leu Thr Asp Met Gly Ser Phe Asp
          1                      5                      10
aga aga gaa agc tgagaacttc ggaaaaggct catctgtcac cctggaraag      558
Arg Arg Glu Ser
          15
ggaaactgta cttttccctg tgaggaaacg gctttgtatt ttctctgtaa taaaatgggg      618
cttctttgga aaaaaaaaaa aa      640

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<210> 258  
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 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 110..304  
 <221> sig\_peptide  
 <222> 110..193  
 <223> Von Heijne matrix  
       score 4.59999990463257  
       seq PLQWSLLVAVVAG/SV  
 <221> polyA\_signal  
 <222> 708..713  
 <221> polyA\_site  
 <222> 732..743

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<400> 258
acttcgcgct gcgcctgcgc agcvcagctc cshgagccct gccaaccatg gtgaacttgg      60
gtctgtcccg ggtggacgac gccgtggctg ccaagcaccg ggcaccggc atg gcc ttt      118
                               Met Ala Phe
ggc ttg cag atg ttc att cag agg aag ttt cca tac cct ttg cag tgg      166
Gly Leu Gln Met Phe Ile Gln Arg Lys Phe Pro Tyr Pro Leu Gln Trp
-25                      -20                      -15                      -10
agc ctc cta gtg gcc gtg gtt gca ggc tct gtg gtc agc tac ggg gtg      214
Ser Leu Leu Val Ala Val Val Ala Gly Ser Val Val Ser Tyr Gly Val
          -5                      1                      5
acg aga gtg gag tgc gag aaa tgc aac aac ctc tgg ctc ttc ctg gag      262
Thr Arg Val Glu Ser Glu Lys Cys Asn Asn Leu Trp Leu Phe Leu Glu
          10                      15                      20
acc gga cag ctc ccc aaa gac agg agc aca gat cag ara agc      304
Thr Gly Gln Leu Pro Lys Asp Arg Ser Thr Asp Gln Xaa Ser
          25                      30                      35
taggagagct ccagcagggg cacagargat tgggggcagg argartctgg aacacakcct      364
tcatgcccc tgaccccagg ccgaccctcc ccacacccta gggtaaccca gtcgtatcct      424
ctgtccgcat gtgtggccag gcctgacaaa cmcctgcaga tggctgtctg cccaacctgg      484
gacctgccca ggaggttggg gcagaaaggg ctctccctgg ggtggtgttt ctctctagg      544
gtattgggat gcatgttctg cactgccagc agagaggggt tgtctggggg ccaccaccta      604
tgggacacgg ggtcgaaggg gcctgtacac tctgtcattt cttttctagc ccctgcatct      664

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ccaacaagtc caaggtgaca gctgggtgcta ggggcgtggg gttaataaat ggcttatcct 724  
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<210> 259  
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<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 201..419

<221> sig\_peptide  
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<223> Von Heijne matrix  
score 6.40000009536743  
seq LSYLPLWLGPWP/CS

<221> polyA\_signal  
<222> 601..606

<221> polyA\_site  
<222> 627..637

<400> 259  
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cctgggtacct gaaaaggtag ctagggtctag gcccttcttc cctttccctt cctctccct 120  
accccagaac tttgggtccc ttcccttct ctctctggta gctccaggag gcctgtgatc 180  
cagctccctg cctagcatcc atg acc tgt tgg atg tta cct cca atc agt ttc 233  
Met Thr Cys Trp Met Leu Pro Pro Ile Ser Phe  
-20 -15  
ctg tcc tac ctg cct ctt tgg ctt gga cct ata tgg cca tgc tct ggc 281  
Leu Ser Tyr Leu Pro Leu Trp Leu Gly Pro Ile Trp Pro Cys Ser Gly  
-10 -5 1  
tct acc ctt ggg aag cct gat ccc ggt gtg tgg ccc agc ttg ttc agg 329  
Ser Thr Leu Gly Lys Pro Asp Pro Gly Val Trp Pro Ser Leu Phe Arg  
5 10 15  
ccc tgg gat gct gca tct cca ggc aac tat gca ctt tcc cgg gga rar 377  
Pro Trp Asp Ala Ala Ser Pro Gly Asn Tyr Ala Leu Ser Arg Gly Xaa  
20 25 30 35  
aac cak tat gav aak tgg ggg cag ggc aca cat tca tct ttg 419  
Asn Xaa Tyr Xaa Xaa Trp Gly Gln Gly Thr His Ser Ser Leu  
40 45  
targaaggtc tggcctgggg tcrngtgaag gagggccag gtcagtctg gggccccagt 479  
gacctgcttt gccattctcc tgggtccgct gctgctccct gtttctggag ctggatgttc 539  
cccacctggc agttgagctg cctgagccaa tgtgtctgtc tttggtaact gagtgaacca 599  
taataaaggg gaacatttgg ccctgtgaaa aaaaaaaa 637

<210> 260  
<211> 1315  
<212> DNA  
<213> Homo sapiens

<220>

<221> CDS

<222> 123..302

<221> sig\_peptide

<222> 123..176

<223> Von Heijne matrix

score 4.30000019073486

seq WTCLKSFPSPTSS/HA

<221> polyA\_signal

<222> 1279..1284

<221> polyA\_site

<222> 1301..1312

<400> 260

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cgccgcagag	ccagtgcaga	atacagaaac	tgcagccatg	accacgcacg	tcaccctgga	120									
ag atg ccc	tgt cca acg	tgg acc tgc	ttg aag agc	ttc ccc tcc	ccg	167									
Met	Pro	Cys	Pro	Thr	Trp	Thr	Cys	Leu	Lys	Ser	Phe	Pro	Ser	Pro	
		-15					-10						-5		
acc agc	agc cat	gca tgc	agc ctc	cac ctt	cct cca	tca tgt	acc agg	215							
Thr	Ser	Ser	His	Ala	Ser	Ser	Leu	His	Leu	Pro	Pro	Ser	Cys	Thr	Arg
		1				5					10				
cta act	ttg aca	caa act	ttg agg	aca gga	atg cat	ttg tca	cgg gca	263							
Leu	Thr	Leu	Thr	Gln	Thr	Leu	Arg	Thr	Gly	Met	His	Leu	Ser	Arg	Ala
		15				20					25				
ttg caa	ggc aca	ttg acc	agg cta	cag tcc	act cca	gca tga	atgarat	312							
Leu	Gln	Gly	Thr	Leu	Thr	Arg	Leu	Gln	Ser	Thr	Pro	Ala			
30		35				40									
gctggaggaa	ggacatgakt	atgcgggtcat	gctgtacacc	tggcgcagct	gttccccgggc	372									
cattccccag	gtgaaatgca	acragcagcc	caaccgakta	raratctatg	araaracagt	432									
araggtgctg	gagccggagg	tcaccaagct	catgaagtcc	atgtattttc	arcgcaaggc	492									
catcgagcgg	ttctgcascg	aggtgaagcg	gctgtgccat	gccgagcgca	ggaaggactt	552									
tgtctctgag	gcctacctcc	tgacccttgg	caagtctatc	aacatgtttg	ctgtcctgga	612									
tgagctaaag	aacatgaast	gcagcgtaaa	raatgaccac	tctgcctaca	agagggcagc	672									
acagttcctg	cggaagatgg	cagatcccca	gtctatccag	gagtcgcaga	acctttccat	732									
gttcctggcc	aaccacaaca	ggatcaccca	gtgtctccac	cagcaacttg	aagtgatccc	792									
aggctatgag	gagctgctgg	ctgacattgt	caacatctgt	gtggattact	acgagaacaa	852									
gatgtacctg	actcccagtg	agaaacatat	gctcctcaag	gtaaaactcc	cctgaggccg	912									
cacccatgga	gcctgggctt	accctctcac	cttcttctta	ttaaaaatcc	gttttaaaaa	972									
acaatgtttc	ttttttctta	aacattgata	cagatcttac	ggcacataat	ggtttgtaac	1032									
ctgttccttt	cctgtaatat	aataaccgt	agtcaccttt	ccagatgtca	ttaaggctat	1092									
ttctacaatg	ttatgtgtaa	tgactgccaa	gtattctgtt	gtattggaac	attgtcatgt	1152									
aacatatccc	ctgtgggttg	atatttgcta	aacttcattg	aacacccttg	tagcagtttt	1212									
tgtgcacatc	tttttgtcaa	ggcaaacttc	ctagaagaga	aattgctggc	tcaaagggaa	1272									
aaacagaata	aatcgttttt	tttatttcaa	aaaaaaaaaa	ccc		1315									

<210> 261

<211> 1035

<212> DNA

<213> Homo sapiens

<220>

<221> CDS  
<222> 98..673

<221> sig\_peptide  
<222> 98..376  
<223> Von Heijne matrix  
score 5.59999990463257  
seq VLLLRQLFAQAEK/WY

<221> polyA\_site  
<222> 1025..1035

<400> 261

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ctcccgttcc tttaggctgc cgccgctgcc tgccgcc atg gca gag ttg ggc cta      115
                                   Met Ala Glu Leu Gly Leu
                                   -90
aat gag cac cat caa aat gaa gtt att aat tat atg cgt ttt gct cgt      163
Asn Glu His His Gln Asn Glu Val Ile Asn Tyr Met Arg Phe Ala Arg
      -85                               -80                               -75
tca aag aga ggc ttg aga ctc aaa act gta gat tcc tgc ttc caa gac      211
Ser Lys Arg Gly Leu Arg Leu Lys Thr Val Asp Ser Cys Phe Gln Asp
      -70                               -65                               -60
ctc aag gag agc agg ctg gtg gag gac acc ttc acc ata gat gaa gtc      259
Leu Lys Glu Ser Arg Leu Val Glu Asp Thr Phe Thr Ile Asp Glu Val
      -55                               -50                               -45                               -40
tct gaa gtc ctc aat gga tta caa gct gtg gtt cat agt gag gtg gaa      307
Ser Glu Val Leu Asn Gly Leu Gln Ala Val Val His Ser Glu Val Glu
      -35                               -30                               -25
tct gag ctc atc aac act gcc tat acc aat gtg tta ctt ctg cga cag      355
Ser Glu Leu Ile Asn Thr Ala Tyr Thr Asn Val Leu Leu Leu Arg Gln
      -20                               -15                               -10
ctg ttt gca caa gct gag aag tgg tat ctt aag cta cag aca gac atc      403
Leu Phe Ala Gln Ala Glu Lys Trp Tyr Leu Lys Leu Gln Thr Asp Ile
      -5                               1                               5
tct gaa ctt gaa aac cga gaa tta tta gaa caa ktt gca gaa ttt gaa      451
Ser Glu Leu Glu Asn Arg Glu Leu Leu Glu Gln Xaa Ala Glu Phe Glu
      10                               15                               20                               25
aaa gca rav att aca tct tca aac aaa aag ccc atc tta dat gtc aca      499
Lys Ala Xaa Ile Thr Ser Ser Asn Lys Lys Pro Ile Leu Xaa Val Thr
      30                               35                               40
aas cca aaa ctt gct cca ctt aat gaa ggt gga aca gca aaa ctc cta      547
Xaa Pro Lys Leu Ala Pro Leu Asn Glu Gly Gly Thr Ala Lys Leu Leu
      45                               50                               55
aac aag gta ata tgt att att ttg aga aac gga aag tct ctc att ctg      595
Asn Lys Val Ile Cys Ile Ile Leu Arg Asn Gly Lys Ser Leu Ile Leu
      60                               65                               70
tcc tgt cat tgc cta ggg tgg aga aac aaa agt gga agg ttt gtt tca      643
Ser Cys His Cys Leu Gly Trp Arg Asn Lys Ser Gly Arg Phe Val Ser
      75                               80                               85
ggt cct ctg agg ata att agt cca ttg cag tagttttact tgatgggtacc      693
Gly Pro Leu Arg Ile Ile Ser Pro Leu Gln
      90                               95
ccatgggcca gaagaggcca tacttaacct tctagagagc ctgaagtagc tctgatcac      753
accttttcaa ggtaaagtga agagcatgaa attttgga gctttattg atggacattt      813
aaagtttgtg atctgcggta acaaggagaa gggtttttaa gtttataaaa attatttatc      873
aattagccgg gtgtggtggt acgtgcctat agtcagagct actcgggagg ctgaggcagg      933

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agaattgctt gaacccggga ggtggagggt gcagtgcgct gagatcacgc cactgcactc 993  
tagcctgggc gacagagcga gactccatct caaaaaaaaa aa 1035

<210> 262  
<211> 696  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 17..463

<221> sig\_peptide  
<222> 17..232  
<223> Von Heijne matrix  
score 3.79999995231628  
seq LMGLALAVYKCQS/MG

<221> polyA\_signal  
<222> 657..662

<221> polyA\_site  
<222> 684..696

<400> 262

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-70 -65  
atc tcc atc ttc cct act atg atg gtg tgt atg atg gcc tgg cga ccc 100  
Ile Ser Ile Phe Pro Thr Met Met Val Cys Met Met Ala Trp Arg Pro  
-60 -55 -50 -45  
att cag gca ctt atg gcc att tca gcc act ttc aag atg tta gaa agt 148  
Ile Gln Ala Leu Met Ala Ile Ser Ala Thr Phe Lys Met Leu Glu Ser  
-40 -35 -30  
tca agc cag aag ttt ctt cag ggt ttg gtc tat ctc att ggg aac ctg 196  
Ser Ser Gln Lys Phe Leu Gln Gly Leu Val Tyr Leu Ile Gly Asn Leu  
-25 -20 -15  
atg ggt ttg gca ttg gct gtt tac aag tgc cag tcc atg gga ctg tta 244  
Met Gly Leu Ala Leu Ala Val Tyr Lys Cys Gln Ser Met Gly Leu Leu  
-10 -5 1  
cct aca cat gca tcg gat tgg tta gcc ttc att gag ccc cct gag aga 292  
Pro Thr His Ala Ser Asp Trp Leu Ala Phe Ile Glu Pro Pro Glu Arg  
5 10 15 20  
atg gag tca gtg gtg gag gac tgc ttt tgt gaa cat gag aaa gca gcg 340  
Met Glu Ser Val Val Glu Asp Cys Phe Cys Glu His Glu Lys Ala Ala  
25 30 35  
cct ggt ccc tat gta ttt ggg tct tat tta cat cct tct tta agc cca 388  
Pro Gly Pro Tyr Val Phe Gly Ser Tyr Leu His Pro Ser Leu Ser Pro  
40 45 50  
gtg gct cct cag cat act ctt aaa cta atc act tat gtt aaa aaa aac 436  
Val Ala Pro Gln His Thr Leu Lys Leu Ile Thr Tyr Val Lys Lys Asn  
55 60 65  
caa aaa act ctt ttc tcc atg gtg ggg tgacaggtcc taaaaggaca 483  
Gln Lys Thr Leu Phe Ser Met Val Gly  
70 75



atgtgcatat	tacgacaaac	acaaaaaaac	tataccataa	cccagggctg	aaaataatgt	543
aaaaaacttt	atTTTTgttt	ccagtacaga	gcaaaacaac	aacaaaaaaa	cataactatg	603
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taaatggttg	araacaatgc	aaaaaaaaaa	aaa			696

<210> 263  
 <211> 868  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 263..481

<221> sig\_peptide  
 <222> 263..322  
 <223> Von Heijne matrix  
 score 11.1999998092651  
 seq ILVVLMLPLAQA/LD

<221> polyA\_site  
 <222> 858..868

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caggagctcc	gggaggcagg gccggcccca cgtcctctgc gcaccaccct gagttggatc 180
ctctgtgcgc	cacccttgag ttggatccag ggctagctgc tgttgacctc cccactccca 240
cgtgcacctc	ctgcctgcag cc atg acg ccc ctg ctc acc ctg atc ctg gtg 292
	Met Thr Pro Leu Leu Thr Leu Ile Leu Val
	-20 -15
gtc ctc atg ggc tta cct ctg gcc cag gcc ttg gac tgc cac gtg tgt 340	
Val Leu Met Gly Leu Pro Leu Ala Gln Ala Leu Asp Cys His Val Cys	
10 -5 1 5	
gcc tac aac gga gac aac tgc ttc aac ccc atg cgc tgc ccg gct atg 388	
Ala Tyr Asn Gly Asp Asn Cys Phe Asn Pro Met Arg Cys Pro Ala Met	
10 15 20	
gtt gcc tac tgc atg acc acg cgc acc tac tac acc ccc acc agg atg 436	
Val Ala Tyr Cys Met Thr Thr Arg Thr Tyr Tyr Thr Pro Thr Arg Met	
25 30 35	
aag gtc agt aag tcc tgc gtg ccc cgc tgc ttc gar nac tgt gta 481	
Lys Val Ser Lys Ser Cys Val Pro Arg Cys Phe Glu Xaa Cys Val	
40 45 50	
tgatggctac	tccaagcacg cgtccaccac ctctgctgc cagtacgacc tctgcaacgg 541
caccggcett	gccaccccg ccaccctggc cctggccccc atcctccttg ccaccctctg 601
gggtctctct	taaagcccc gaggcagacc cactcaagaa caaagctctc gagacacact 661
gctayacct	ckcaccckc tcaccctgcc tcaccctcca cactccctgc gacctcctca 721
gccatgccca	gggtcaggac tgtgggcaag aagacaccg acctcccca accaccacac 781
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<210> 264  
 <211> 775

<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 42..299

<221> sig\_peptide  
<222> 42..101  
<223> Von Heijne matrix  
score 5.40000009536743  
seq WFWHSSALGLVLA/PP

<221> polyA\_site  
<222> 762..775

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-20  
cca cag tgg ttt gtt cat tca tca gcg tta ggc ttg gtc ctg gct cca 104  
Pro Gln Trp Phe Val His Ser Ser Ala Leu Gly Leu Val Leu Ala Pro  
-15 -10 -5 1  
cct ttc tcc tct ccg ggc act gac ccc acc ttt ccg tgt att tac tgt 152  
Pro Phe Ser Ser Pro Gly Thr Asp Pro Thr Phe Pro Cys Ile Tyr Cys  
5 10 15  
agg cta tta aat atg atc atg acc cgc ctt gca ttt tca ttc atc acc 200  
Arg Leu Leu Asn Met Ile Met Thr Arg Leu Ala Phe Ser Phe Ile Thr  
20 25 30  
tgt tta tgc cca aat tta aag gaa gtt tgt ctc att ttg cca gaa aaa 248  
Cys Leu Cys Pro Asn Leu Lys Glu Val Cys Leu Ile Leu Pro Glu Lys  
35 40 45  
aat tgt aat agt cga cac gct gga ttt gta ggg cca sca aaa ttg cgg 296  
Asn Cys Asn Ser Arg His Ala Gly Phe Val Gly Pro Xaa Lys Leu Arg  
50 55 60 65  
cag tgaaactwkk ttcwcttcta aagcccttca tttcccaaa ggtaagctc 349  
Gln  
tcgaaacccc atttgatcct tggttcctat ttcgatcctc ctttggaatc tgaaaatcgg 409  
tctccatgtt gtatgcaaat taaaaktgtc cttgtttgtt actcttccaa cacagggtat 469  
cagggaraaa gaggccttat ctgttcctcc atccccctg ttttgacaga ctgctaagaa 529  
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tcaggggtag acaagcttgt cctagtgtc tgcttcaggt cttatcagaa gaaaccagg 649  
aatagaaaag gtagatgcct tgacttttgt ccctgttgtg gggactaaag tgttttttgc 709  
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aaaaaa 775

<210> 265  
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<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 198..431

<221> sig\_peptide  
 <222> 198..260  
 <223> Von Heijne matrix  
 score 6.90000009536743  
 seq LLACGSLLPGLWQ/HL

<221> polyA\_site  
 <222> 1064..1074

<400> 265  
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 taagaaacaa gcatacctgt gtaagtgaat tatcttaatt tgtgttgaat caagttagga 180  
 gagacagatt ctcatga atg tgt cct gtg ttc tca aag cag ctg cta gcc 230  
 Met Cys Pro Val Phe Ser Lys Gln Leu Leu Ala  
 -20 -15  
 tgt ggg tct ctc cta cct ggg tta tgg cag cac ctc aca gcc aat cac 278  
 Cys Gly Ser Leu Leu Pro Gly Leu Trp Gln His Leu Thr Ala Asn His  
 -10 -5 1 5  
 tgg cct cca ttc tcc sct ttc ctc tgt aca gtt tgc tct ggt tcc tca 326  
 Trp Pro Pro Phe Ser Xaa Phe Leu Cys Thr Val Cys Ser Gly Ser Ser  
 10 15 20  
 gag cag att tcc gag tat act gct tca gcc acg ccc cca ctg tgc cgt 374  
 Glu Gln Ile Ser Glu Tyr Thr Ala Ser Ala Thr Pro Pro Leu Cys Arg  
 25 30 35  
 tcc ctg aac caa gag cca ttc gty tca aga gcc att cgt cca aag tac 422  
 Ser Leu Asn Gln Glu Pro Phe Val Ser Arg Ala Ile Arg Pro Lys Tyr  
 40 45 50  
 tct atc acc tagcattgt akccatacca agccgggctt cctacttccc 471  
 Ser Ile Thr  
 55  
 tctgctcccc ttggtttcct cctgtraart aaatctcact gacccttgat gcasctccaa 531  
 gcatatataa tatatatata ataaaaccat abtctaaaaa attcaaacca ggawaaataa 591  
 asccaraaat ttgtatggga aaaatctgca caaatttatt tggccagcat gggtatcatg 651  
 gctctattga atttatcctt gaccgtcttt aaagccaaag caaacgggat aaagtgatca 711  
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 aaaacaattc ttakctgggc gcggtggctc wcacctgtar tccaacact ttgggaggcc 831  
 gaggtgggag gatcatgagg tggggagatc aamaccatcc tggctaacat ggtgaaaccc 891  
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 ctactcggga ggctgaggca agagaatggg gtgaacccca gggggcggag cctgcagtga 1011  
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 aaah 1075

<210> 266  
 <211> 981  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 279..473

<221> sig\_peptide  
 <222> 279..362  
 <223> Von Heijne matrix

score 4.40000009536743

seq SCFLVALIIWCYL/RE

<221> polyA\_signal

<222> 944..949

<221> polyA\_site

<222> 970..981

<400> 266

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cgctgaggct cataggtgg gcttcccgaa gccttcaccc gttgcccggt tcccgggatc      180
gggcccaccc tgccgcccag gaagaggacg accctgaccg cccattgag tttcctcca      240
gcaaagccaa ccctcaccgc tggtcggtgg gccatacc atg gga aag gga cat cag      296
                               Met Gly Lys Gly His Gln
                               -25
cgg ccc tgg tgg aag gtg ctg ccc ctc agc tgc ttc ctc gtg gcg ctg      344
Arg Pro Trp Trp Lys Val Leu Pro Leu Ser Cys Phe Leu Val Ala Leu
      -20                      -15                      -10
atc atc tgg tgc tac ctg agg gag gag agc gag gcg gac cag tgg ttg      392
Ile Ile Trp Cys Tyr Leu Arg Glu Glu Ser Glu Ala Asp Gln Trp Leu
      -5                      1                      5                      10
aga cag gtg tgg gga gag gtg cca gag ccc agt gat cgt tct gag gag      440
Arg Gln Val Trp Gly Glu Val Pro Glu Pro Ser Asp Arg Ser Glu Glu
      15                      20                      25
cct gag act cca gct gcc tac aga gcg aga act tgacgggggtg cccgctgggg      493
Pro Glu Thr Pro Ala Ala Tyr Arg Ala Arg Thr
      30                      35
ctggcaggaa gggagccgac asccgccctt cggatttgat ktcacgtttg cccgtgactg      553
tcctggctat gcktgcgctc tcagcactra argacttggc tgggtggatgg ggcacttggc      613
tatgctgatt cgcgtgaagg cggavcaaaa tctcagcaaa tcggaaactg ctctcscct      673
ggctcttgat ktccaaggat tccatcggca aaacttctca ratccttggg gaaggtttca      733
gttgccactgt atgctgttgg atttgccaag tctttgtata acataatcat gtttccaaag      793
cacttctggt gacacttgtc atccagtgtt agtttgcagg taatttgctt tctgagatag      853
aatatctggc agaagtgtga aactgtattg catgtgcgg cctgtgcaag gaacacttcc      913
acatgtgagt tttacacaac aacaaatgaa aataaatttt aattttataa tatgggaaaa      973
aaaaaaaaa                                     981
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<210> 267

<211> 1031

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 12..644

<221> sig\_peptide

<222> 12..92

<223> Von Heijne matrix

score 4

seq LTFFSGVYGTCIG/AT

<221> polyA\_signal

<222> 1002..1007

<221> polyA\_site

<222> 1020..1031

<400> 267

acaccaagga g atg ctc ctt ctt agt att aca act gct tat aca ggt ctg	50
Met Leu Leu Leu Ser Ile Thr Thr Ala Tyr Thr Gly Leu	
-25 -20 -15	
gaa tta act ttc ttc tct ggt gta tat gga acc tgt att ggt gct aca	98
Glu Leu Thr Phe Phe Ser Gly Val Tyr Gly Thr Cys Ile Gly Ala Thr	
-10 -5 1	
aat aaa ttt gga gca gaa gag ara agc ctt att gga ctt tct ggc att	146
Asn Lys Phe Gly Ala Glu Glu Xaa Ser Leu Ile Gly Leu Ser Gly Ile	
5 10 15	
ttc atc ggc att gga gaa att tta ggt gga agc ctc ttc ggc ctg ctg	194
Phe Ile Gly Ile Gly Glu Ile Leu Gly Gly Ser Leu Phe Gly Leu Leu	
20 25 30	
agc aag aac aat cgt ttt ggt aga aat cca gtt gtg ctg ttg ggc atc	242
Ser Lys Asn Asn Arg Phe Gly Arg Asn Pro Val Val Leu Leu Gly Ile	
35 40 45 50	
ctg gtg cac ttc ata gct ttt tat cta ata ttt ctc aac atg cct gga	290
Leu Val His Phe Ile Ala Phe Tyr Leu Ile Phe Leu Asn Met Pro Gly	
55 60 65	
gat gcc ccg att gct cct gtt aaa gga act gac agc agt gct tac atc	338
Asp Ala Pro Ile Ala Pro Val Lys Gly Thr Asp Ser Ser Ala Tyr Ile	
70 75 80	
aaa tcc agc aaa raa ttt gcc att ctc tgc akt ttt ctg tkg ggc ctt	386
Lys Ser Ser Lys Xaa Phe Ala Ile Leu Cys Xaa Phe Leu Xaa Gly Leu	
85 90 95	
gga aac agc tgc ttt aat acc cas ctg ctt akt atc tkg ggc ttt ctg	434
Gly Asn Ser Cys Phe Asn Thr Xaa Leu Leu Xaa Ile Xaa Gly Phe Leu	
100 105 110	
tat tct gaa rac agc gcc cca kca ttt gcc atc ttc aat ttt gtt cag	482
Tyr Ser Glu Xaa Ser Ala Pro Xaa Phe Ala Ile Phe Asn Phe Val Gln	
115 120 125 130	
tct att tgc gca gcc gtg gca ttt ttc tac agc aac tac ctt ctc ctt	530
Ser Ile Cys Ala Ala Val Ala Phe Phe Tyr Ser Asn Tyr Leu Leu Leu	
135 140 145	
cac tgg caa ctc ctg gtc atg gtk atw ttt ggg ttt ttk gga aca att	578
His Trp Gln Leu Leu Val Met Val Ile Phe Gly Phe Xaa Gly Thr Ile	
150 155 160	
tct ttc ttc act gtg gaa tgg gaa sct gcc gcc ttt gta scc cgc ggc	626
Ser Phe Phe Thr Val Glu Trp Glu Xaa Ala Ala Phe Val Xaa Arg Gly	
165 170 175	
tct gac tac cga agt atc tgatctgggtg tccgtgaggg gacacgtatg	674
Ser Asp Tyr Arg Ser Ile	
180	
acctcagaaa cacagctgga cacagagctt ggtggaagaa gtcgcctttg atcttcacta	734
tatattgggt gatgttcagt atggaaaatc aagggtattaa gactgttaaa tcagccagag	794
tkggtgttca agttttacaga tatgagttat ttaaagcaag tagaataagg gaaagctgtt	854
ctgtcaactg taattgttca aagatgttgt ttttcatttc atctatctca attcttataa	914
tcattgttata gaatgtaaat gttttcttct ctctcctgct cttgtttggaa gatcctgcct	974
tgatttagaa tactaggcca tatgtcatat aaatattttt tctggaaaaa aaaaaaa	1031

<210> 268  
<211> 1283  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 91..459

<221> sig\_peptide  
<222> 91..330  
<223> Von Heijne matrix  
score 7.69999980926514  
seq LVLFLSLALLVTP/TS

<221> polyA\_site  
<222> 1271..1281

<400> 268  
tattccttgg agttccacga ctgaattaag actgttgtgg grdccataat tttcaaatac 60  
ttgccctata ttcgtgttga ggggttcacac atg agc aca tgg tat ttg gca ctt 114  
Met Ser Thr Trp Tyr Leu Ala Leu  
-80 -75  
aat aag tcc tat aag aat aaa gac agc gtt agg att tat ctc agc ttg 162  
Asn Lys Ser Tyr Lys Asn Lys Asp Ser Val Arg Ile Tyr Leu Ser Leu  
-70 -65 -60  
tgc aca gtg agc att aaa ttt aca tac ttt cat gat ata cag act aat 210  
Cys Thr Val Ser Ile Lys Phe Thr Tyr Phe His Asp Ile Gln Thr Asn  
-55 -50 -45  
tgt ctt aca aca tgg aaa cat tcg aga tgc aga ttt tat tgg gca ttt 258  
Cys Leu Thr Thr Trp Lys His Ser Arg Cys Arg Phe Tyr Trp Ala Phe  
-40 -35 -30 -25  
ggg ggt tcc att tta cag cac tca gtg gat ccc ctt gtt ttg ttc cta 306  
Gly Gly Ser Ile Leu Gln His Ser Val Asp Pro Leu Val Leu Phe Leu  
-20 -15 -10  
agc ctg gcc ctg tta gtg aca ccc act tcc acc cct tct gct aar ata 354  
Ser Leu Ala Leu Leu Val Thr Pro Thr Ser Thr Pro Ser Ala Lys Ile  
-5 1 5  
car agc ctt caa att gac ctc cct gga ggc tgg agg ctg gcc act gac 402  
Gln Ser Leu Gln Ile Asp Leu Pro Gly Gly Trp Arg Leu Ala Thr Asp  
10 15 20  
agg atc ttt acc ctc tcc ccc gta ccc atg gac rgc ccc ctc atc ctt 450  
Arg Ile Phe Thr Leu Ser Pro Val Pro Met Asp Xaa Pro Leu Ile Leu  
25 30 35 40  
cat cag ttg taaaggtaga tatttgttcc ttggagtcca acatcatgct 499  
His Gln Leu  
gttcagaata taatgagatc aatagttgaa aaactagata tacatgccac ccwgacaaag 559  
ctattaagtt attaagtgtc agccctggat cttggccttat tgtgaaatgt taattatttt 619  
atcactcyat taagaagctg tgggctccat ctcagcattg aaaagggact aatttgcctc 679  
gtttttggaat tgaattagct ttcaggccas cagggcactg tttggtaaat tgcctttttcc 739  
agtactagca tgtttttctcc ctccatagcc tctgttagct tctgagcttg taacctccag 799  
ggaaavatga gaataattcac ccttttaata tgtgtagaga ccatgcaaga ccattgtctt 859  
ctaataatta gaaatactta gccagattct ctatagtaaa cccggagatt gggagggctg 919  
ctttctactt ggtgcatcct tctgcgcttc taatgatttt taaaaatctg ttaataattg 979  
atgtttttctg gctgggcaca gtggctcacg cctgtaatcc cagcactttg ggaggccaag 1039  
gagggcagat catgaggtca ggagattgar accatcctgg ctaacacggg gaaaccccg 1099  
ctctactaaa aatacaaaar aattakccgg gcatggtagt gggcgctgt gtaccagct 1159

actggggagg ctgaggcarg araatcgctt gaacctggga ggcggaggtt gcastragct 1219  
gagatggtgc caccgcactc tagcctgggt gacagagcga gacttcattt caaaaaaaaaa 1279  
aamc 1283

<210> 269  
<211> 1777  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 70..327

<221> sig\_peptide  
<222> 70..147  
<223> Von Heijne matrix  
score 9.60000038146973  
seq WLIALASWSWALC/RI

<221> polyA\_signal  
<222> 1741..1746

<221> polyA\_site  
<222> 1763..1774

<400> 269  
agccccggttt cgtgccccgcg gccgactgcg casctgtccg cgagtctgag atacttacag 60  
agagctaca atg gaa aag tcc tgg atg ctg tgg aac ttt gtt gaa aga tgg 111  
Met Glu Lys Ser Trp Met Leu Trp Asn Phe Val Glu Arg Trp  
-25 -20 -15  
cta ata gcc ttg gct tca tgg tct tgg gct ctc tgc cgt att tct ctt 159  
Leu Ile Ala Leu Ala Ser Trp Ser Trp Ala Leu Cys Arg Ile Ser Leu  
-10 -5 1  
tta cct tta ata gtg act ttt cat ctg tat gga ggc att atc tta ctt 207  
Leu Pro Leu Ile Val Thr Phe His Leu Tyr Gly Gly Ile Ile Leu Leu  
5 10 15 20  
ttg tta ata ttc ata tca atw kca ggt att ctg tat aaa ttc cas gat 255  
Leu Leu Ile Phe Ile Ser Ile Xaa Gly Ile Leu Tyr Lys Phe Xaa Asp  
25 30 35  
gta ttg ctt tat ttt ccw kaa cag yya tcc tct tca cgt ctt tat gat 303  
Val Leu Leu Tyr Phe Pro Xaa Gln Xaa Ser Ser Ser Arg Leu Tyr Asp  
40 45 50  
tcc cat gcc cac tgg cmt tcg rca taaaaaatt ttcatacagaa ccaaagatgg 357  
Ser His Ala His Trp Xaa Ser Xaa  
55 60  
aatacgtctg aatottatatt tgatacgata cactggagac aattcaccct attccccaac 417  
tataatattat tttcatggga atgcaggcaa cataggtcac aggttggcca aatgcattac 477  
ttatgtttggt taacctcaaa gttaaccttt tgctggttga ttatcgagga tatggaaaaa 537  
gtgaaggaga agcaagtga gaaggactct acttagattc tgaagctgtg ttagactacg 597  
tgatgactag acctgacctt gataaaacaa aaatttttct ttttggccgt tccttgggtg 657  
garcagtggc tattcatttg gcttctgaaa attcacatag gatttcagcc attatggtgg 717  
agaacacatt ttttaagcata ccacatatgg ccagcacttt attttcattc tttccgatgc 777  
gttaccttcc tttatggtgc tacaaaaata aatttttgtc ctacagaaaa atctctcagt 837  
gtagaatgcc ttcacttttc atctctggac tctcagatca attaattcca ccagtaatga 897  
tgaaacaact ttatgaactc tccccatctc ggactaagan attagccatt tttccagatg 957

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ggactcacia tgacacatgg cagtgccaa gctatttcac tgcacttgaa cagttcatca 1017
aagaagtcgt aaagagccat tctcctgaag aaatggcaaa aacttcatct aatgtaacaa 1077
ttatataatg tttccctttt tgattattgc attgtatttt aatttggtgca gaatgataaa 1137
gaatgttcct tttagaagtg tgttatgtct gtacctgtct gaagagtgc attaaacttt 1197
gaaaggactt cactgctcct ttacgatatt ccaaatagtt ttttacattg gaaaaactaa 1257
ttcttgggat tctttcatac attttcatca aaactttcag tgtgattatg tattcatatc 1317
ttcagtttaa tatgtcagta taatagatat tgttcaaaag tttcttggtg ctaaagtggg 1377
gtaatctgtt acacagatga atagctagat gtggaaagag atatgtaaac aagaaacctt 1437
tgggtattgt ttcttaagta aatattggga caatcatggt aagcaaactt agttctgtaa 1497
ctgcattttt cactttaaaa gttaaataaa atgcatgatg gtattttatt ccttgaatta 1557
tgcaatgcaa cattttacat gtaaataagca ctggatcatat actgatgtat atggttatct 1617
gggttatatc tatttttatg taaactctat ttttgttttt ggcaagaagt gaaattgaga 1677
cttatgtgca ggttgccatt gaattttgct ctggatgaatg ctgagatcca gctttttctt 1737
acaaataaat gggaccctgt tttccaaaaa aaaaaamcm 1777
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<210> 270

<211> 970

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 12..497

<221> sig\_peptide

<222> 12..104

<223> Von Heijne matrix

score 5.5

seq LVGVLFVSVTTG/PW

<221> polyA\_signal

<222> 935..940

<221> polyA\_site

<222> 955..967

<400> 270

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aggtctccaa g atg gcg gcc gcc tgg ccg tct ggt ccg kct gct ccg gag 50
          Met Ala Ala Ala Trp Pro Ser Gly Pro Xaa Ala Pro Glu
          -30          -25          -20
gcc gtg acg gcc aga ctc gtt ggt gtc ctg tgg ttc gtc tca gtc act 98
Ala Val Thr Ala Arg Leu Val Gly Val Leu Trp Phe Val Ser Val Thr
          -15          -10          -5
aca gga ccc tgg ggg gct gtt gcc acc tcc gcc ggg ggc gag gag tcg 146
Thr Gly Pro Trp Gly Ala Val Ala Thr Ser Ala Gly Gly Glu Glu Ser
          1          5          10
ctt aag tgc gag gac ctc aaa gtg gga caa tat att tgt aaa gat cca 194
Leu Lys Cys Glu Asp Leu Lys Val Gly Gln Tyr Ile Cys Lys Asp Pro
          15          20          25          30
aaa ata aat gac gct acg caa gaa cca gtt aac tgt aca aac tac aca 242
Lys Ile Asn Asp Ala Thr Gln Glu Pro Val Asn Cys Thr Asn Tyr Thr
          35          40          45
gct cat gtt tcc tgt ttt cca gca ccc aac ata act tgt aag gat tcc 290
Ala His Val Ser Cys Phe Pro Ala Pro Asn Ile Thr Cys Lys Asp Ser
          50          55          60
```



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agt ggc aat gaa aca cat ttt act ggg aac gaa gtt ggt ttt ttc aag      338
Ser Gly Asn Glu Thr His Phe Thr Gly Asn Glu Val Gly Phe Phe Lys
      65                      70                      75
ccc ata tct tgc cga aat gta aat ggc tat tcc tac aat gag cag tcg      386
Pro Ile Ser Cys Arg Asn Val Asn Gly Tyr Ser Tyr Asn Glu Gln Ser
      80                      85                      90
cat gtc tct ttt tct tgg atg gtt ggg agc aga tcg att tta cct tgg      434
His Val Ser Phe Ser Trp Met Val Gly Ser Arg Ser Ile Leu Pro Trp
      95                      100                      105                      110
ata ccc tgc ttt ggg ttt gtt aaa btt tyg cac tgt agg gtt tkg tgg      482
Ile Pro Cys Phe Gly Phe Val Lys Xaa Xaa His Cys Arg Val Xaa Trp
      115                      120                      125
aat tgg gag cct aat tgatttcaty cttatttcaa tgcagattgt tggaccttca      537
Asn Trp Glu Pro Asn
      130
aatggaagta gttacattat agattactat ggaaccagac ttacaagact gagtattact      597
aatgaaacat ttagaaaaac gcaattatat ccataaatat tttttaaaaag aaacagattt      657
gagcctcctt gattttaata gagaacttct agtgtatgga tttaaagatt tctctttttc      717
attcatatac cattttatga gttctgtata attttttgtg gtttttgttt tgttgagtta      777
aagtatatta ttgtgagatt tatttaatag gacttccttt gaaagctgta taatagtgtt      837
tctcgggctt ctgtctctat gagagatagc ttattactct gatactcttt aatcttttac      897
aaaggcaagt tgccacttgt catttttggt tctgaaaaat aaaagtataa cttattcaca      957
aaaaaaaaaa mms                                                    970
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<210> 271
<211> 645
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 90..383
<221> sig_peptide
<222> 90..200
<223> Von Heijne matrix
      score 4.90000009536743
      seq MLIMLGIFFNVHS/AV
```

```
<221> polyA_signal
<222> 609..614
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<221> polyA_site
<222> 632..643
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```
<400> 271
atctctgccc ccctgcgagg gcatactggg ctttctccca ccgctttccg agcccgttg      60
cacctoggcg atccccgact cccttcttt atg gcg tcg ctc ctg tgc tgt ggg      113
                        Met Ala Ser Leu Leu Cys Cys Gly
                        -35                      -30
ccg aag ctg gcc gcc tgc ggc atc gtc ctc agc gcc tgg gga gtg atc      161
Pro Lys Leu Ala Ala Cys Gly Ile Val Leu Ser Ala Trp Gly Val Ile
      -25                      -20                      -15
atg ttg ata atg ctc gga ata ttt ttc aat gtc cat tcc gct gtg ttg      209
Met Leu Ile Met Leu Gly Ile Phe Phe Asn Val His Ser Ala Val Leu
```

```

      -10      -5      1
att gag gac gtt ccc ttc acg gag aaa gat ttt gag aac ggc ccc car      257
Ile Glu Asp Val Pro Phe Thr Glu Lys Asp Phe Glu Asn Gly Pro Gln
      5      10      15
aac ata tac aac ctt tac rag caa ktc agc tac aac tgt ttc atc gct      305
Asn Ile Tyr Asn Leu Tyr Xaa Gln Xaa Ser Tyr Asn Cys Phe Ile Ala
      20      25      30      35
gca ggc ctt tac ctc ctc ctc gga ggc ttc tct ttc tgc caa ktt cgg      353
Ala Gly Leu Tyr Leu Leu Leu Gly Gly Phe Ser Phe Cys Gln Xaa Arg
      40      45      50
ctc aat aag cgc aag gaa tac atg gtg cgc tagggccccg gcgcgtttcc      403
Leu Asn Lys Arg Lys Glu Tyr Met Val Arg
      55      60
ccgctccagc ccctcctcta tttaaaract ccctgcacgc tktcacccag gtgcgcgtccc      463
acccttgccg ggcgcctctg tgggactggg tttcccgggc rararactga atcccttctc      523
ccatctctgg catccggccc ccgtggarar ggctgaggct ggggggctgt tccgtctctc      583
cacccttcgc tgtgtcccg atctcaataa agagaatctg ctctcttcaa aaaaaaaaaa      643
my                                                                    645
```

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<210> 272
<211> 773
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 332..541
<221> sig_peptide
<222> 332..376
<223> Von Heijne matrix
      score 3.59999990463257
      seq FLPCCLLWSVFNP/ES
<221> polyA_signal
<222> 739..744
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<221> polyA_site
<222> 761..773
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<400> 272
aaaacaattc atgcctttca tagtttatta ttattaaagt ctaaacaaaa ttgcaatttc      60
ttaggtaacc ttatatctac aataaatgaa gattaccctc aaatgctaga agctgtctag      120
gtccgctcgg tgtgtcagat tttcctcaga ttagatgtgc caataaccaa gtttattcag      180
taaacaactt gtacttgttt catctggttt tattactctc acccataaac agtaatgact      240
ctctgaccct ctggaaatat gtaatgcttc caatcttgct ttgtgtatct catttaattt      300
gttataaggt agtactgatt ttagcatatt a atg cga ttt ctt cct tgt tgt      352
                                Met Arg Phe Leu Pro Cys Cys
                                -15      -10
ttg ctt tgg tct gtg ttc aat cca gag agc tta aat tgt cat tat ttt      400
Leu Leu Trp Ser Val Phe Asn Pro Glu Ser Leu Asn Cys His Tyr Phe
      -5      1      5
ghk ndd gaa amc tgt att ttt gyt agt tta caa tat tat gaa att tca      448
Xaa Xaa Glu Xaa Cys Ile Phe Xaa Ser Leu Gln Tyr Tyr Glu Ile Ser
      10      15      20
```

```

ctt cag gag aaa ctg ctg ggc ttc ctg tgg ctt tgt ttt ctt agt tac      496
Leu Gln Glu Lys Leu Leu Gly Phe Leu Trp Leu Cys Phe Leu Ser Tyr
25          30          35          40
ttt ttc cgt gcc gtg tat ttt tta att gat ttt tct tct ttt act      541
Phe Phe Arg Ala Val Tyr Phe Leu Ile Asp Phe Ser Ser Phe Thr
          45          50          55
tgaaaagaaa gtgtttttatt ttcaaactctg gtccatattt acatttctagt tcagagccaa      601
gccttaaact gtacagaatt tccactgtaa ttaaaactat ttagtgtag ttataaatag      661
ccttcaaaaa gagagattct ccattacacg atcacctgca tcacagccca tggatgaatgt      721
atgtttctgc atagcgaaat aaaaatggca aatgcactga aaaaaaaaaa aa      773

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<210> 273  
 <211> 566  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 43..222

<221> sig\_peptide  
 <222> 43..177  
 <223> Von Heijne matrix  
 score 4  
 seq ENFLSLLSKSCSA/DP

<221> polyA\_signal  
 <222> 530..535

<221> polyA\_site  
 <222> 555..566

```

<400> 273
aacgagtgga ggtgtggcta gtggctgtga tgagataaat cc atg cat agc ctt      54
                                   Met His Ser Leu
                                   -45
ttc att gcg agc ttg aaa gtt ctt ttc tat tac agt ttt agc ttt agg      102
Phe Ile Ala Ser Leu Lys Val Leu Phe Tyr Tyr Ser Phe Ser Phe Arg
-40          -35          -30
ttt aat tgg ttc gac tgc ctt ctc cac aat ttg ggc gag aat ttc ctt      150
Phe Asn Trp Phe Asp Cys Leu Leu His Asn Leu Gly Glu Asn Phe Leu
-25          -20          -15          -10
agc ctt ctc agc aaa agt tgt tct gcg gac ccg tct ggg tca act ttc      198
Ser Leu Leu Ser Lys Ser Cys Ser Ala Asp Pro Ser Gly Ser Thr Phe
          -5          1          5
atg agg gac att gag aca aac aaa tgaaatatgg gttaaagtac tctgagcagc      252
Met Arg Asp Ile Glu Thr Asn Lys
10          15
tacaaaaaga araccagtct atcctgtctgg agacagtggc cacgtgaara aagagctctt      312
gcagtatgaa agaccacatg gaaagagagg ccacatggaa ccaacagtca gcatcttggt      372
ttcggacacg tgaaraaatt catctcarac tgtgtatcct aaatcaggca cttgctgaat      432
ctaactacac gagtgagacc agttgacaac acatggagca racatgagct gttctcagtg      492
artcctacac aaattcctga ctcaaacac tgtgagcaat aaaatggttg ttattttaag      552
ccaaaaaaaa aaaa      566

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<210> 274  
<211> 455  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 115..231

<221> sig\_peptide  
<222> 115..180  
<223> Von Heijne matrix  
score 5  
seq HLFVTWSSQRALS/HP

<221> polyA\_signal  
<222> 419..424

<221> polyA\_site  
<222> 445..455

<400> 274  
aacctgccag tkatgcaaat gccaaaatgt gggatcatcat atagtatatatt tgaaaccttt 60  
ctgaacatgt acaccaccca atgctagagg ctgacttgga aaccggtggg tgca atg 117  
Met  
ccc gag gct gtg gaa caa tca gcc cat ctc ttt gtg acc tgg agc agt 165  
Pro Glu Ala Val Glu Gln Ser Ala His Leu Phe Val Thr Trp Ser Ser  
-20 -15 -10  
cag agg gcc ctc agt cac ccc gcc cca ttc ctc acc ara raa aar aat 213  
Gln Arg Ala Leu Ser His Pro Ala Pro Phe Leu Thr Xaa Xaa Lys Asn  
5 1 5 10  
cca ttt cta tgg aag ctc tgacgtaact tcagtgtttt ctacaataact 261  
Pro Phe Leu Trp Lys Leu  
15  
ctctctgccc cgcgccatta aaacagttct tttgttaaaa aatavcctaa tgggtccaact 321  
ctgtgtgtctg ttcttccaaa tgtttataat acacattatt tataaatatg tctgtttggg 381  
aagctaagaa caagctagtt tttacaacac aaatggaaat aaatgcaatt attataaaaa 441  
tycaaaaaaa aaaa 455

<210> 275  
<211> 673  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 232..384

<221> sig\_peptide  
<222> 232..300  
<223> Von Heijne matrix  
score 3.70000004768372  
seq FFLCAAFPLGAGV/KM

<221> polyA\_signal  
<222> 650..655

<221> polyA\_site  
<222> 662..673

<400> 275  
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ctgaaggacc cagcttaggt tcttccactt aggcctcaat tcccttcctt ttccaggggc 120  
agccttagtt tcccatggcc ctgaaacaca cacatttccc ccttcctttc ccagaagcca 180  
ctggccccc atagcaccca gtgcacacct tttacaagtg gaagaactag g atg gct 237  
Met Ala  
ttc caa agt ctt cta gaa atg aag ttc ttt ctc tgt gca gct ttc ccc 285  
Phe Gln Ser Leu Leu Glu Met Lys Phe Phe Leu Cys Ala Ala Phe Pro  
-20 -15 -10  
ctt gga gca gga gtg aag atg ttt cat tat ctt ggg cct ggg aaa cca 333  
Leu Gly Ala Gly Val Lys Met Phe His Tyr Leu Gly Pro Gly Lys Pro  
-5 1 5 10  
ctt cyy cag gct tct ccc tcc ccc cac ccc cat agg amc agg att tgg 381  
Leu Xaa Gln Ala Ser Pro Ser Pro His Pro His Arg Xaa Arg Ile Trp  
15 20 25  
cct tagcttctgg gcctatcsgc tgccttccct cttyttccta ccacctcttc 434  
Pro  
tgccttccct trawctctgt tgggcttggg gatcttagtt ttcttttgtt tatttcccat 494  
ctcatttttt tcttctgggc agttttttta aggggggggtg ttgtgggttt ttgtttttgt 554  
tttgcttctg aaaaarcatt tgcctttctt cctctcccaa cataacaatc gtggtaacag 614  
aatgcgactg ctgatttacc gatgtattta atgtaagtaa aaaaaggaaa aaaaraaaa 673

<210> 276  
<211> 639  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 143..427

<221> sig\_peptide  
<222> 143..286  
<223> Von Heijne matrix  
score 7.5  
seq FVILLLLFIFTVVS/LV

<221> polyA\_signal  
<222> 606..611

<221> polyA\_site  
<222> 628..639

<400> 276  
aatcgcttca gcagcaccct ctgagacaag agccactatt tctgattcag atcacctgtc 60  
atcgaagttt aaagaagggg aaacaggaga cagaaatata ctgaaccaa aagattcaaa 120  
agagcaagtg gaatctctaa ga atg gct tcc agc cac tgg aat gaa acc act 172  
Met Ala Ser Ser His Trp Asn Glu Thr Thr

															-45				-40		
acc	tct	gtt	tat	cag	tac	ctt	ggt	ttt	caa	gtt	caa	aaa	att	tac	cct	220					
Thr	Ser	Val	Tyr	Gln	Tyr	Leu	Gly	Phe	Gln	Val	Gln	Lys	Ile	Tyr	Pro						
			-35				-30				-25										
ttc	cat	gac	aac	tgg	aac	act	gcc	tgc	ttt	gtc	atc	ctg	ctt	tta	ttt	268					
Phe	His	Asp	Asn	Trp	Asn	Thr	Ala	Cys	Phe	Val	Ile	Leu	Leu	Leu	Phe						
			-20				-15				-10										
ata	ttt	aca	gtg	gta	tct	tta	gtg	gtg	ctg	gct	ttc	ctt	tat	gaa	gtg	316					
Ile	Phe	Thr	Val	Val	Ser	Leu	Val	Val	Leu	Ala	Phe	Leu	Tyr	Glu	Val						
			-5				1				5				10						
ctt	gam	wgc	tgc	tgc	tgt	gta	aaa	aac	aaa	acc	gtg	aaa	gac	ttg	aaa	364					
Leu	Xaa	Xaa	Cys	Cys	Cys	Val	Lys	Asn	Lys	Thr	Val	Lys	Asp	Leu	Lys						
			15				20				25										
agt	gaa	ccc	aac	cct	ctt	ara	akt	atg	atg	gac	aac	atc	aga	aaa	cgt	412					
Ser	Glu	Pro	Asn	Pro	Leu	Xaa	Xaa	Met	Met	Asp	Asn	Ile	Arg	Lys	Arg						
			30				35				40										
gaa	act	gaa	gtg	gtc	taacactcta			taraaaaatga			acaaaatctc			tgaaagcagc		467					
Glu	Thr	Glu	Val	Val																	
			45																		

tcaacctctt	ctgaraaaaa	aaatatattc	tgaggccaac	tgttgtctaca	aaacaaattc	527
tgactgaatg	gttaaaacat	ttctagtara	aggggaaaaa	aaakttaaac	atgcactggt	587
tggtgtgata	scattttcat	taaatataca	gtaaaactyc	aaaaaaaaaa	aa	639

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<210> 277
<211> 772
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 284..463

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<221> sig_peptide
<222> 284..379
<223> Von Heijne matrix
score 3.79999995231628
seq TFINITLWLGLSC/OR
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<221> polyA_site
<222> 762..772
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<400> 277															
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cctgtattgc	acttttgggt	ttaaggactg	gacccagagt	tcctgaaagc	caaactccat										120
aagctgctca	gtaagttcca	agcacatagc	cggtckhggg	atgcgattcg	gtcgaggtct										180
gttgaatgaa	ggtagacgca	gcaggcagtt	tgtccttacc	agtgacctgg	aagacggtgg										240
cacttcctga	gtgagctcac	ttaccttccc	tgaatggtga	ggc atg gat	gaa tat										295
				Met	Asp	Glu	Tyr								
							-30								
tcc tgg tgg tgc cac gtg tta gag gtg gta aag ggt caa atg ttt act															343
Ser Trp Trp Cys His Val Leu Glu Val Val Lys Gly Gln Met Phe Thr															
	-25					-20				-15					
ttt att aat att aca tta tgg ctt ggt tct ctg tgt cag cga ttt ttc															391
Phe Ile Asn Ile Thr Leu Trp Leu Gly Ser Leu Cys Gln Arg Phe Phe															
	-10					-5				1					

```
tat gcc tcg ggt act tat ttc cta ata tat atc agc aca gta acg cct 439
Tyr Ala Ser Gly Thr Tyr Phe Leu Ile Tyr Ile Ser Thr Val Thr Pro
5          10          15          20
agc tgg agg ctt tgt ctt gtt agt tgataaatta gtggtaacag gtagatttgg 493
Ser Trp Arg Leu Cys Leu Val Ser
25
ttacctccca aagtgtctggg attrcagacg tgagccaccg cgcctggccg aaacaattct 553
tttgaaagag agaagtctcc ctgtgtttgcg caggctggtc tcagactcct ggggtcaagt 613
gagcctcctg ctttcgcctc ctaaagtgtc gggattacag gcgtgagcca ccgcacccgg 673
acagatgtgt tgattttaaa gtgggtatga ggcctgagcc ctggagtttg agaccagcct 733
ggacaacatg gcaagaccct gtctctccaa aaaaaaaaa 772
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<210> 278

<211> 840

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 162..671

<221> sig\_peptide

<222> 162..398

<223> Von Heijne matrix

score 4.09999990463257

seq QGVLFICFTCARS/FP

<221> polyA\_signal

<222> 805..810

<221> polyA\_site

<222> 830..840

<400> 278

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aaaaactgag gcctgggagc aggaacctgt aggcagcgct tgagggtagc gggatagcag 60
ctgcaacgcg cgtgggaggc gggggctctg ggcggaacaa aaatcacagg atgtcagagg 120
atgtttcccg ggaagaactg ggataaaggg gtcccagcac c atg gag gac ccg aac 176
Met Glu Asp Pro Asn
-75
```

```
cct gaa gag aac atg aag cag cag gat tca ccc aag gag aga agt ccc 224
Pro Glu Glu Asn Met Lys Gln Gln Asp Ser Pro Lys Glu Arg Ser Pro
-70 -65 -60
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cag agc cca gga ggc aac atc tgc cac ctg ggg gcc ccg aag tgc acc 272
Gln Ser Pro Gly Gly Asn Ile Cys His Leu Gly Ala Pro Lys Cys Thr
-55 -50 -45
```

```
cgc tgc ctc atc acc ttc gca gat tcc aag ttc cag gag cgt cac atg 320
Arg Cys Leu Ile Thr Phe Ala Asp Ser Lys Phe Gln Glu Arg His Met
-40 -35 -30
```

```
aag cgg gag cac cca gcg gac ttc gtg gcc cag aag ctg cag ggg gtc 368
Lys Arg Glu His Pro Ala Asp Phe Val Ala Gln Lys Leu Gln Gly Val
-25 -20 -15
```

```
ctc ttc atc tgc ttc acc tgc gcc cgc tcc ttc ccc tcc tcc aaa gcc 416
Leu Phe Ile Cys Phe Thr Cys Ala Arg Ser Phe Pro Ser Ser Lys Ala
-10 -5 1 5
```

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ckr rkc acc cac car cgc agc cac ggt cca rcc gcc aag ccc acc ctg 464
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Xaa	Xaa	Thr	His	Gln	Arg	Ser	His	Gly	Pro	Xaa	Ala	Lys	Pro	Thr	Leu	
		10						15				20				
ccg	gtt	gca	acc	act	act	gcc	car	ccc	acc	ttc	cct	tgt	cct	gac	tgt	512
Pro	Val	Ala	Thr	Thr	Thr	Ala	Gln	Pro	Thr	Phe	Pro	Cys	Pro	Asp	Cys	
		25					30				35					
ggc	aaa	acc	ttt	ggg	cag	gct	gtt	tct	ctg	arg	cgg	cac	csc	caa	atr	560
Gly	Lys	Thr	Phe	Gly	Gln	Ala	Val	Ser	Leu	Xaa	Arg	His	Xaa	Gln	Xaa	
		40				45				50						
cat	gar	gtc	cgt	gcc	cct	cct	ggc	acc	ttc	gcc	tgc	aca	rad	tgc	ggt	608
His	Glu	Val	Arg	Ala	Pro	Pro	Gly	Thr	Phe	Ala	Cys	Thr	Xaa	Cys	Gly	
55					60				65					70		
cag	gac	ttt	gct	car	gaa	rca	ggg	ctg	cat	caa	cac	tac	att	cgg	cat	656
Gln	Asp	Phe	Ala	Gln	Glu	Xaa	Gly	Leu	His	Gln	His	Tyr	Ile	Arg	His	
			75				80					85				
gcc	cgg	ggg	gga	ctc	tgagttcagc	ttaagcctct	ccacggtgac	gggtggctct								711
Ala	Arg	Gly	Gly	Leu												
			90													
gtggctggta	ggactcaccc	atgatatggg	gtgcaggaac	tctggggggcc	ctgaaggatt											771
tgcttcctc	ccctgggaag	gcagagggt	cttaataaag	aggacccaka	agattcttaa											831
aaaaaaaa																840

210 211 212 213 220 221 222 221 222 223 221 222 221 222 400

<210> 279  
 <211> 840  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 63..632

<221> sig\_peptide  
 <222> 63..308  
 <223> Von Heijne matrix  
 score 4.40000009536743  
 seq NLPHLQVVGLTWG/HI

<221> polyA\_signal  
 <222> 808..813

<221> polyA\_site  
 <222> 829..840

<400> 279	
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ga atg tat gtd tgg ccc tgt gct gtg gtc ctg gcc cag tac ctt tgg	107
Met Tyr Val Trp Pro Cys Ala Val Val Leu Ala Gln Tyr Leu Trp	
-80 -75 -70	
ttt cac aga aga tct ctg cca ggc aag gcc atc tta gag att gga gct	155
Phe His Arg Arg Ser Leu Pro Gly Lys Ala Ile Leu Glu Ile Gly Ala	
-65 -60 -55	
gga gtg agc ctt cca gga att ttg gct gcc aaa tgt ggt gca gaa gta	203
Gly Val Ser Leu Pro Gly Ile Leu Ala Ala Lys Cys Gly Ala Glu Val	
-50 -45 -40	
ata ctg tca gac agc tca gaa ctg cct cac tgt ctg gaa gtc tgt cgg	251
Ile Leu Ser Asp Ser Ser Glu Leu Pro His Cys Leu Glu Val Cys Arg	



-35	-30	-25	-20	
caa agc tgc caa atg aat aac ctg cca cat ctg cag gtg gta gga cta				299
Gln Ser Cys Gln Met Asn Asn Leu Pro His Leu Gln Val Val Gly Leu				
	-15	-10	-5	
aca tgg ggt cat ata tct tgg gat ctt ctg gct cta cca cca caa gat				347
Thr Trp Gly His Ile Ser Trp Asp Leu Leu Ala Leu Pro Pro Gln Asp				
	1	5	10	
att atc ctt gca tct gat gtg ttc ttt gaa cca gaa rat ttt gaa gac				395
Ile Ile Leu Ala Ser Asp Val Phe Phe Glu Pro Glu Xaa Phe Glu Asp				
	15	20	25	
att ttg gct aca ata tat ttt ttg atg cac aar aat ccc aag gtc caa				443
Ile Leu Ala Thr Ile Tyr Phe Leu Met His Lys Asn Pro Lys Val Gln				
	30	35	40	45
ttg tgg tct act tat caa gtt agg art gct gac tgg tca ctt gaa gct				491
Leu Trp Ser Thr Tyr Gln Val Arg Xaa Ala Asp Trp Ser Leu Glu Ala				
	50	55	60	
tta ctc tac aaa tgg gat atg aaa tgt gtc cac att cct ctt gag tct				539
Leu Leu Tyr Lys Trp Asp Met Lys Cys Val His Ile Pro Leu Glu Ser				
	65	70	75	
ttt gat gca gac aaa gaa rat ata gca gaa tct acc ctt cca gga aga				587
Phe Asp Ala Asp Lys Glu Xaa Ile Ala Glu Ser Thr Leu Pro Gly Arg				
	80	85	90	
Cat aca gtt gaa atg ctg gtc att tcc ttt gca aag gac agt ctc				632
His Thr Val Glu Met Leu Val Ile Ser Phe Ala Lys Asp Ser Leu				
	95	100	105	
tgaattatac ctacaacctg ttctgggaca gtatcaatac tgatgagcaa cctgggcacac				692
aaactatgag cagaccactt cagcttgaga atgcagtggg tctgaagatg gtcaagtctg				752
tttgccttar attttgatgt cacctagaca acacttaaac tcatatgaaa caaaaattaa				812
ataacgtatt acaagcaaaa aaaaaaaa				840

<210> 280

<211> 849

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 21..362

<221> sig\_peptide

<222> 21..200

<223> Von Heijne matrix

score 4.80000019073486

seq LVILSLKSQTLDA/ET

<221> polyA\_signal

<222> 821..826

<221> polyA\_site

<222> 838..849

<400> 280

agtaagtccc cccgcctcgc atg atg gct gcg gtg ccg ccg ggc ctg gag ccg

Met Met Ala Ala Val Pro Pro Gly Leu Glu Pro

-60 -55 -50



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tcc ccc cag gcc ctg gag gac tcg ggc ccg gtg aat atc tca gtc tca 101
Ser Pro Gln Ala Leu Glu Asp Ser Gly Pro Val Asn Ile Ser Val Ser
-95 -90 -85
atc acc cta acc ctg gac cca ctg aaa ccc ttc gga ggg tat tcc cgc 149
Ile Thr Leu Thr Leu Asp Pro Leu Lys Pro Phe Gly Gly Tyr Ser Arg
-80 -75 -70
aac gtc acc cat ctg tac tca acc atc tta ggg cat cag att gga ctt 197
Asn Val Thr His Leu Tyr Ser Thr Ile Leu Gly His Gln Ile Gly Leu
-65 -60 -55 -50
tca ggc agg gaa gcc cac gag gag ata aac atc acc ttc acc ctg cct 245
Ser Gly Arg Glu Ala His Glu Glu Ile Asn Ile Thr Phe Thr Leu Pro
-45 -40 -35
aca gcg tgg agc tca gat gac tgc gcc ctc cac ggt cac tgt gag cag 293
Thr Ala Trp Ser Ser Asp Asp Cys Ala Leu His Gly His Cys Glu Gln
-30 -25 -20
gtg gta ttc aca gcc tgc atg acc ctc acg gcc agc cct ggg gtg ttc 341
Val Val Phe Thr Ala Cys Met Thr Leu Thr Ala Ser Pro Gly Val Phe
-15 -10 -5
ccg tca ctg tac agc cac cgc act gtg ttc ctg aca cgt aca gca acg 389
Pro Ser Leu Tyr Ser His Arg Thr Val Phe Leu Thr Arg Thr Ala Thr
1 5 10 15
cca cgc tct ggt aca aga tct tca caa ctg cca gag atg cca aca caa 437
Pro Arg Ser Gly Thr Arg Ser Ser Gln Leu Pro Glu Met Pro Thr Gln
20 25 30
aat acg ccc aaa att aca atc ctt tct ggt gtt ata agg ggg cca ttg 485
Asn Thr Pro Lys Ile Thr Ile Leu Ser Gly Val Ile Arg Gly Pro Leu
35 40 45
gaa aag tct atc atg ctt taaatcccaa gcttacagtg attgttccag 533
Glu Lys Ser Ile Met Leu
50
atgatgaccg ttcattaata aatttgcac tcacgcacac cagttacttc ctctttgtga 593
tggtgataac aatgttttgc tatgctgtta tcaagggcag acctagcaaa ttgcgtcaga 653
gcaatcctga attttgtccc gagaagggtg ctttggctga agcctaattc cacagctcct 713
tggttttttga gagagactga gagaaccata atccttgccg gctgaaccca gcctgggcct 773
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acctgatggg acgtttccac gtgtctctag agaaggatcc tggatctagc tggtcacgac 1013
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<210> 282  
 <211> 671  
 <212> DNA  
 <213> Homo sapiens

<220>  
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 <222> 1..201  
 <221> sig\_peptide

<222> 1..63

<223> Von Heijne matrix  
score 5.09999990463257  
seq LLLKIWLLQRPES/QE

<221> polyA\_signal

<222> 637..642

<221> polyA\_site

<222> 660..671

<400> 282

atg ctg gga ggt gac cat agg gct ctg ctt tta aag ata tgg ctg ctt	48
Met Leu Gly Gly Asp His Arg Ala Leu Leu Leu Lys Ile Trp Leu Leu	
-20 -15 -10	
caa agg cca gag tca cag gaa gga ctt ctt cca ggg aga tta gtg gtg	96
Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro Gly Arg Leu Val Val	
-5 1 5 10	
atg gag agg aga gtt aaa aat gac ctc atg tcc ttc ttg tcc acg gtt	144
Met Glu Arg Arg Val Lys Asn Asp Leu Met Ser Phe Leu Ser Thr Val	
15 20 25	
ttg ttg agt ttt cac tct tct aat gca agg gtc tca cac tgt gaa cca	192
Leu Leu Ser Phe His Ser Ser Asn Ala Arg Val Ser His Cys Glu Pro	
30 35 40	
ctt agg atg tgatcacttt caggtggcca ggaatgttga atgtctttgg	241
Leu Arg Met	
45	
ctcagttcat ttaaaaaaga tatctatttg aaagttctca rarttgtaca tatgtttcac	301
agtacaggat ctgtacataa aagtttcttt cctaaaccat tcaccaagag ccaatatcta	361
ggcattttct tggtagcaca aattttctta ttgcttaraa aattgtcctc cttgttattt	421
ctgtttgtaa racttaagtg agttaggtct ttaaggaaag caacgctcct ctgaaatgct	481
tgtctttttt ctgttgccga aatarctggg ccttttttcgg gagttaratg tatarartgt	541
ttgtatgtaa acatttcttg taggcatacac catgaacaaa gatataattt ctattttattt	601
attatatgtg cacttcaaga agtcactgtc agagaaataa agaattgtct taaatgtcaa	661
aaaaaaaaaa	671

<210> 283

<211> 1601

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 39..1034

<221> sig\_peptide

<222> 39..134

<223> Von Heijne matrix  
score 6.09999990463257  
seq LPLLTSA LHGLQQ/QH

<221> polyA\_signal

<222> 1566..1571

<221> polyA\_site

<222> 1587..1597

<400> 283

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                               Met Ile Xaa Leu Arg Asp
                               -30
aca gct gcc tcc ctc cgc ctt gag aga gac aca agg cag ttg cca ctg      104
Thr Ala Ala Ser Leu Arg Leu Glu Arg Asp Thr Arg Gln Leu Pro Leu
-25                               -20                               -15
ctc acc agt gcc ctg cac gga ctg cag cag cag cac cca gcc ttc tct      152
Leu Thr Ser Ala Leu His Gly Leu Gln Gln Gln His Pro Ala Phe Ser
-10                               -5                               1                               5
ggg gtg gca cgg ctg gcc aag cgg tgg gtg cgt gcc cag ctt ctt ggt      200
Gly Val Ala Arg Leu Ala Lys Arg Trp Val Arg Ala Gln Leu Leu Gly
10                               15                               20
gag ggt ttc gct gat gag agc ctg gat ctg gtg gcc gct gcc ctt ttc      248
Glu Gly Phe Ala Asp Glu Ser Leu Asp Leu Val Ala Ala Ala Leu Phe
25                               30                               35
ctg cac cct gag ccc ttc acc cct ccg agt tcc ccc cag gtt ggc ttc      296
Leu His Pro Glu Pro Phe Thr Pro Pro Ser Ser Pro Gln Val Gly Phe
40                               45                               50
ctt cga ttc ctt ttc ttg gta tca acg ttt gat tgg aag aac aac ccc      344
Leu Arg Phe Leu Phe Leu Val Ser Thr Phe Asp Trp Lys Asn Asn Pro
55                               60                               65                               70
ctc ttt gtc aac ctc aat aat gag ctc act gtg gag gag cag gtg gar      392
Leu Phe Val Asn Leu Asn Asn Glu Leu Thr Val Glu Glu Gln Val Glu
75                               80                               85
atc cgc agt ggc ttc ctg gca gct cgg gca cag ctc ccc gtc atg gtc      440
Ile Arg Ser Gly Phe Leu Ala Ala Arg Ala Gln Leu Pro Val Met Val
90                               95                               100
att gtt acc ccc caa rac cgc aaa aac tct gtg tgg aca cag gat gga      488
Ile Val Thr Pro Gln Xaa Arg Lys Asn Ser Val Trp Thr Gln Asp Gly
105                               110                               115
ccc tca gcc car atc ctg cag cag ctt gtg gtc ctg gca gct gaa scc      536
Pro Ser Ala Gln Ile Leu Gln Gln Leu Val Val Leu Ala Ala Glu Xaa
120                               125                               130
ctg ccc atg tta rar aas cag ctc atg gat ccc cgg gga cct ggg gac      584
Leu Pro Met Leu Xaa Xaa Gln Leu Met Asp Pro Arg Gly Pro Gly Asp
135                               140                               145                               150
atc agg aca gkg ttc cgg ccg ccc ttg gac att tac gac gtg ctg att      632
Ile Arg Thr Xaa Phe Arg Pro Pro Leu Asp Ile Tyr Asp Val Leu Ile
155                               160                               165
cgc ctg tct cct cgc cat atc ccg cgg cac cgc cag gct gtg gac tcr      680
Arg Leu Ser Pro Arg His Ile Pro Arg His Arg Gln Ala Val Asp Ser
170                               175                               180
cca gct gcc tcc ttc tgc cgg ggc ctg ctc agc cag ccg ggg ccc tca      728
Pro Ala Ala Ser Phe Cys Arg Gly Leu Leu Ser Gln Pro Gly Pro Ser
185                               190                               195
tcc ctg atg ccc gtg ctg ggc tak gat cct cct cag ctc tat ctg acg      776
Ser Leu Met Pro Val Leu Gly Xaa Asp Pro Pro Gln Leu Tyr Leu Thr
200                               205                               210
cag ctc arg gag gcc ttt ggg gat ctg gcc ctt ttc ttc tat gac cag      824
Gln Leu Xaa Glu Ala Phe Gly Asp Leu Ala Leu Phe Phe Tyr Asp Gln
215                               220                               225                               230
cat ggt gga gag gtg att ggt gtc ctc tgg aag ccc acc agc ttc cag      872
His Gly Gly Glu Val Ile Gly Val Leu Trp Lys Pro Thr Ser Phe Gln
235                               240                               245

```

cgc	cag	ccc	ttc	aag	gcc	tcc	agc	aca	aag	ggg	cgc	atg	gtg	atg	tct	920
Pro	Gln	Pro	Phe	Lys	Ala	Ser	Ser	Thr	Lys	Gly	Arg	Met	Val	Met	Ser	
			250					255					260			
cga	ggt	ggg	gag	cta	gta	atg	gtg	ccc	aat	gtt	gaa	gca	atc	ctg	gag	968
Arg	Gly	Gly	Glu	Leu	Val	Met	Val	Pro	Asn	Val	Glu	Ala	Ile	Leu	Glu	
			265					270					275			
gac	ttt	gct	gtg	ctg	ggt	gaa	ggc	ctg	gtg	cag	act	gtg	gag	gcc	cga	1016
Asp	Phe	Ala	Val	Leu	Gly	Glu	Gly	Leu	Val	Gln	Thr	Val	Glu	Ala	Arg	
			280					285					290			
agt	gag	agg	tgg	act	gtg	tgatcccagc tctggagcaa gctgtagacg										1064
Ser	Glu	Arg	Trp	Thr	Val											
			295			300										
gacagcagga	cattggacct					ctagagcaag	atgtcagtag					gatgacctcc	accctccttg			1124
gacatgaatc	ctccatggag					ggcctgctgg	ctgaacatgc					tgaatcatct	ccaacaaaac			1184
ccagccccc	ctttctctct					gatgctccag	cattggggca					ggggcatggt	ggcccatgta			1244
gtctcctggg	cctcaccatc					ccagaagagg	agtgggagcc					agctcagaga	aggaactgaa			1304
cccaggagat	ccatccacct					attagccctg	ggcctggacc					tccttgcgat	ttcccactcc			1364
tttcttagtc	ttcttccaga					aacagagaag	gggagtgtgt					ctctgggagag	gctctgtctc			1424
cttctgtctg	ccaggacctg					tgctagact	tagcatgcc					ttcatgcag	tgtcaggcct			1484
ttagatggga	cccagcgaaa					atgtggccct	tctgagtcac					atcacgcaca	ctgagcagtg			1544
gaaaggggct	atatgtgtat					gaatagacca	cattgaagga					qcaaaaaaaaa	aaamcch			1601

Year	1950	1951	1952	1953	1954	1955	1956	1957	1958	1959	1960	1961	1962	1963	1964	1965	1966	1967	1968	1969	1970	1971	1972	1973	1974	1975	1976	1977	1978	1979	1980	1981	1982	1983	1984	1985	1986	1987	1988	1989	1990	1991	1992	1993	1994	1995	1996	1997	1998	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022	2023	2024	2025	2026	2027	2028	2029	2030	2031	2032	2033	2034	2035	2036	2037	2038	2039	2040	2041	2042	2043	2044	2045	2046	2047	2048	2049	2050	2051	2052	2053	2054	2055	2056	2057	2058	2059	2060	2061	2062	2063	2064	2065	2066	2067	2068	2069	2070	2071	2072	2073	2074	2075	2076	2077	2078	2079	2080	2081	2082	2083	2084	2085	2086	2087	2088	2089	2090	2091	2092	2093	2094	2095	2096	2097	2098	2099	2100
1950	1951	1952	1953	1954	1955	1956	1957	1958	1959	1960	1961	1962	1963	1964	1965	1966	1967	1968	1969	1970	1971	1972	1973	1974	1975	1976	1977	1978	1979	1980	1981	1982	1983	1984	1985	1986	1987	1988	1989	1990	1991	1992	1993	1994	1995	1996	1997	1998	1999	2000	2001	2002	2003	2004	2005	2006	2007	2008	2009	2010	2011	2012	2013	2014	2015	2016	2017	2018	2019	2020	2021	2022	2023	2024	2025	2026	2027	2028	2029	2030	2031	2032	2033	2034	2035	2036	2037	2038	2039	2040	2041	2042	2043	2044	2045	2046	2047	2048	2049	2050	2051	2052	2053	2054	2055	2056	2057	2058	2059	2060	2061	2062	2063	2064	2065	2066	2067	2068	2069	2070	2071	2072	2073	2074	2075	2076	2077	2078	2079	2080	2081	2082	2083	2084	2085	2086	2087	2088	2089	2090	2091	2092	2093	2094	2095	2096	2097	2098	2099	2100	

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<210> 284
<211> 1206
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 69..263

<221> sig_peptide
<222> 69..125
<223> Von Heijne matrix
score 3.90000009536743
seq ALSMSSFSFHSSS/CS
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<221> polyA_signal
<222> 1173..1178
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```
<221> polyA_site
<222> 1196..1205
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<400> 284																		
acatttgtga ctttaccaat accctcccag ttcttgatag acagctgtag gttgctgggt																		60
tcaagaat atg ggt ggg ata tgg aat gct ctt tca atg tct agc ttc agt																		110
Met Gly Gly Ile Trp Asn Ala Leu Ser Met Ser Ser Phe Ser																		
-15 -10																		
ttt cat tca tcc tcc tgc tca gca ctg tca gcc aag agc tta ctc agc																		158
Phe His Ser Ser Ser Cys Ser Ala Leu Ser Ala Lys Ser Leu Leu Ser																		
-5 1 5 10																		
aga cac cac ata ctg cag cag ttc cta gtg aga aaa tct gtg cca cta																		206
Arg His His Ile Leu Gln Gln Phe Leu Val Arg Lys Ser Val Pro Leu																		
15 20 25																		
gaa aat gct tca ctt cca ttt cct cac ctg qqc aqt tct ctg ttt aaa																		254

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<210> 285
<211> 536
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 115..285

<221> sig_peptide
<222> 115..204
<223> Von Heijne matrix
      score 3.70000004768372
      seq SMLLLTVYGGYLC/SV

<221> polyA_signal
<222> 505..510

<221> polyA_site
<222> 525..536

<400> 285
acgagtgctg cgttcggctg tgctgggaag ttgcgtagac agtggcctcg agaccctgcc      60
tgcttgagga ggccctcggtt ggatgcgaag gagctgcagc atccagggga caag atg      117
                                         Met
                                         -30
cca act ggc aag cag cta gct gac att ggc tat aag acc ttc tct acc      165
Pro Thr Gly Lys Gln Leu Ala Asp Ile Gly Tyr Lys Thr Phe Ser Thr
          -25                      -20                      -15
tcc atg atg ctt ctc act gtg tat ggg ggg tac ctc tgc agt gtc cga      213
Ser Met Met Leu Leu Thr Val Tyr Gly Gly Tyr Leu Cys Ser Val Arg
          -10                      -5                      1
gtc tac cac tat ttc cag tgg cgc agg gcc caa cgc caa gcc qca qaa      261

```

```
Val Tyr His Tyr Phe Gln Trp Arg Arg Ala Gln Arg Gln Ala Ala Glu
  5              10              15
gaa cag aag dac tca gga atc atg tagaactggg gggctttttc tcctgagcar 315
Glu Gln Lys Xaa Ser Gly Ile Met
20              25
asakgcccac ggcattgctgt ggagagactt cacctgccac catttccagg tcaacaggac 375
tagagcgttg atggtttttca aaccctgttg gaagaaagtg cccatggttt ctctggttct 435
gccartttga cagtttatgg argcttttga atcgtaatar caatgtgagg gtgargtaca 495
cctacagaca ttaaataatt tgctgtgtca aaaaaaaaaa a 536
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<210> 286  
<211> 529  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 90..344

<221> sig\_peptide  
<222> 90..140  
<223> Von Heijne matrix  
score 8.19999980926514  
seq LLLITAILAVAVG/FP

<221> polyA\_signal  
<222> 500..505  
  
<221> polyA\_site  
<222> 515..527

<400> 286  
aatatrarac agctacaata ttccagggcc artcacttgc catttctcat aacagcgtca 60  
gagagaaaga actgactgar acgtttgag atg aag aaa gtt ctc ctc ctg atc 113  
Met Lys Lys Val Leu Leu Leu Ile  
-15 -10  
aca gcc atc ttg gca gtg gct gtw ggt ttc cca gtc tct caa gac cag 161  
Thr Ala Ile Leu Ala Val Ala Val Gly Phe Pro Val Ser Gln Asp Gln  
-5 1 5  
gaa cga gaa aaa aga agt atc agt gac agc gat gaa tta gct tca ggr 209  
Glu Arg Glu Lys Arg Ser Ile Ser Asp Ser Asp Glu Leu Ala Ser Gly  
10 15 20  
wtt ttt gtg ttc cct tac cca tat cca ttt cgc cca ctt cca cca att 257  
Xaa Phe Val Phe Pro Tyr Pro Tyr Pro Phe Arg Pro Leu Pro Pro Ile  
25 30 35  
cca ttt cca aga ttt cca tgg ttt aga cgt aat ttt cct att cca ata 305  
Pro Phe Pro Arg Phe Pro Trp Phe Arg Arg Asn Phe Pro Ile Pro Ile  
40 45 50 55  
cct gaa tct gcc cct aca act ccc ctt cct agc gaa aag taaacaaraa 354  
Pro Glu Ser Ala Pro Thr Thr Pro Leu Pro Ser Glu Lys  
60 65  
ggaaaagtca crataaacct ggtcacctga aattgaaatt gagccacttc cttgaaraat 414  
caaaattcct gttaataaaa raaaaacaaa tgtaattgaa atagcacaca gcattctcta 474  
gtcaatatct ttagtgatct tctttaataa acatgaaagc aaaaaaaaaa aaacc 529



<210> 287  
<211> 493  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 57..311

<221> sig\_peptide  
<222> 57..107  
<223> Von Heijne matrix  
score 8.19999980926514  
seq LLLITAILAVAVG/FP

<221> polyA\_signal  
<222> 467..472

<221> polyA\_site  
<222> 482..493

400> 287  
aacttgccat ttctcataac agcgtcagag agaaagaact gactgaaacg tttgag atg 59  
Met  
aag aaa gtt ctc ctc ctg atc aca gcc atc ttg gca gtg gct gtt ggt 107  
Lys Lys Val Leu Leu Leu Ile Thr Ala Ile Leu Ala Val Ala Val Gly  
-15 -10 -5  
ttc cca gtc tct caa gac cak gaa cga gaa aaa aga agt atc agt gac 155  
Phe Pro Val Ser Gln Asp Xaa Glu Arg Glu Lys Arg Ser Ile Ser Asp  
5 10 15  
agc gat gaa tta gct tca ggg ttt ttt gtg ttc cct tac cca tat cca 203  
Ser Asp Glu Leu Ala Ser Gly Phe Phe Val Phe Pro Tyr Pro Tyr Pro  
20 25 30  
ttt cgc cca ctt cca cca att cca ttt cca aga ttt cca tgg ttt aga 251  
Phe Arg Pro Leu Pro Pro Ile Pro Phe Pro Arg Phe Pro Trp Phe Arg  
35 40 45  
cgt aat ttt cct att cca ata cct gaa tct gcc cct aca act ccc ctt 299  
Arg Asn Phe Pro Ile Pro Ile Pro Glu Ser Ala Pro Thr Thr Pro Leu  
50 55 60  
ccg agc gaa aag taaacaagaa ggaaaagtca cgataaacct ggtcacctga 351  
Pro Ser Glu Lys  
65  
aattgaaatt gagccacttc cttgargaat caaaattcct gttaataaaa gaaaaacaaa 411  
tgtaattgaa atagcacaca gcattctcta gtcaatatct ttagtgatct tctttaataa 471  
acatgaaagc aaaaaaaaaa aa 493

<210> 288  
<211> 521  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS

<222> 96..302

<221> sig\_peptide

<222> 96..182

<223> Von Heijne matrix

score 5

seq ELSLLPSSLWVLA/TS

<221> polyA\_site

<222> 501..514

<400> 288

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aagagacgtc accggctgcg cccttcagta tcgcggacgg aagatggcgt ccgccaccgc 60
tctcatocag cggctgcgga actgggcgtc cgggc atg acc tgc agg gga agc 113
                               Met Thr Cys Arg Gly Ser
                               -25
tgc agc tac gct acc agg aga tct cca agc gaa ctc agc ctc ctc cca 161
Cys Ser Tyr Ala Thr Arg Arg Ser Pro Ser Glu Leu Ser Leu Leu Pro
          -20          -15          -10
agc tcc ctg tgg gtc cta gcc aca agc tct cca aca att act att gca 209
Ser Ser Leu Trp Val Leu Ala Thr Ser Ser Pro Thr Ile Thr Ile Ala
          -5          1          5
ctc gcg atg gcc gcc ggg aat ctg tgc ccc ctt cca tca tca tkt cgt 257
Leu Ala Met Ala Ala Gly Asn Leu Cys Pro Leu Pro Ser Ser Xaa Arg
10          15          20          25
cgc aaa agg cgc tgg tgt cag gca asc car caa ara gct ctg ctg 302
Xaa Lys Arg Arg Trp Cys Gln Ala Xaa Gln Gln Xaa Ala Leu Leu
          30          35          40
tagctgccac tgaaaaraag gcggtgactc cagctcctcc cataaagagg tgggagctgt 362
cctcggacca gccttacctg tgacactgca ccctcacggc caccgacta ctttgctctc 422
ttggatttcc tccagggaga atgtgacctt atttatgaca aatacgtara gctcaggtat 482
cacttctagt ttacttttaa aaaataaaaa aatagagac 521

```

<210> 289

<211> 811

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 161..526

<221> sig\_peptide

<222> 161..328

<223> Von Heijne matrix

score 4.19999980926514

seq XSPLLTLALLGQC/SL

<221> polyA\_site

<222> 799..811

<400> 289

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aaaaaattgc agtgctgaag aacttgacc cgcaaaaggc tgtccctccc aaacctggga 60
ttctgggctc actgagttca cctgcgagtc agcctacct gactgctct ggtctagtac 120
aaacaggctg ctggcattga ggtctgctac aaaaanarta atg gtc cca tgg ccc 175

```

```

Met Val Pro Trp Pro
-55
agg ggc aag gtg aaa act gct cct att ccc atc tct agg ttt cct ttc 223
Arg Gly Lys Val Lys Thr Ala Pro Ile Pro Ile Ser Arg Phe Pro Phe
-50 -45 -40
ctc cct acc cac gac cca ccc acc cca gca cat tgg tct cca gca tct 271
Leu Pro Thr His Asp Pro Pro Thr Pro Ala His Trp Ser Pro Ala Ser
-35 -30 -25 -20
cat cag cag ttt aaa cat kkg tca ccc ctc ctc act ttg gcc ctg ctg 319
His Gln Gln Phe Lys His Xaa Ser Pro Leu Leu Thr Leu Ala Leu Leu
-15 -10 -5
ggg cag tgc tct ctg ttc arc aat ttg agg aaa aaa ctt gca ggg caa 367
Gly Gln Cys Ser Leu Phe Xaa Asn Leu Arg Lys Lys Leu Ala Gly Gln
1 5 10
aaa gca aaa aaa tta cct tcc ttc tcc agc ctg ccc ctg aca ctc tgg 415
Lys Ala Lys Lys Leu Pro Ser Phe Ser Ser Leu Pro Leu Thr Leu Trp
15 20 25
cca tta act cct caa ttt gct gag ctc act aca gtg gca caa aaa aaa 463
Pro Leu Thr Pro Gln Phe Ala Glu Leu Thr Val Ala Gln Lys Lys
30 35 40 45
ttg agg tgg tcc ggg acc cta ggt tgg ggt cca gtt ccc agc tgg gtt 511
Leu Arg Trp Ser Gly Thr Leu Gly Trp Gly Pro Val Pro Ser Trp Val
50 55 60
Caa ttt ttt tta ggg tgaatggagg garagttggg gactgaaaas ccttcaaara 566
Gln Phe Phe Leu Gly
65
caatgttatt acagcaktct ccccttatcc aaaktttcct tttcctgadt ttcagtttagc 626
tatgggtcaac cgcttggaaa atakttgaac acagtacaat aaratatttt gaggtctggga 686
cttgggtggctc atgcctgtaa taatcccagg actttgtgar accaaktttg aaggatcact 746
tgaacccagg aktttgarac caccctgggc aacatrgtra gacctcatct ctacaaaaaa 806
aaaaa 811

<210> 290
<211> 625
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 210..332

<221> sig_peptide
<222> 210..299
<223> Von Heijne matrix
score 8.10000038146973
seq ITCLLAFWVPASC/IQ

<221> polyA_signal
<222> 594..599

<221> polyA_site
<222> 613..625

<400> 290
acagggtcsmc ttaacatctc ttgatttgag ccactccac tgatcatcagc tttcacctgg 60

```

```

attatcgtga cagcctccta ctgcttctct atcatgtggc cagagctatc ttccctaata 120
atgcattgca tagttgatca agtcactctc tggcctaata ccttccttgg ctccctgctg 180
ccctcaggat aaagtctgga cccctcagc atg gct tgt gag act cat ggt gtc 233
                               Met Ala Cys Glu Thr His Gly Val
                               -30                -25
ctt gtc cct gct cac ctg tct ggt ctg atc act tgc ctt ctt gca ttc 281
Leu Val Pro Ala His Leu Ser Gly Leu Ile Thr Cys Leu Leu Ala Phe
      -20                -15                -10
tgg gtc cca gcc tcc tgt atc cag aga tgc agt ggc tct cca ttg cca 329
Trp Val Pro Ala Ser Cys Ile Gln Arg Cys Ser Gly Ser Pro Leu Pro
      -5                1                5                10
ctc tgattcctcc tttcttttgg tcacagagaa aggggtacttt ctctgtcaaa 382
Leu
tctcaactta gaattgaatt cctccaagga gctttggcta tactctctcc cwcgaccccc 442
accctggcat actacacara tcactctggg ctactttgcc tgcctaattg tcatctcccc 502
agtaaactgt aagctccttg agggcaagga ttgtgttggg atttttgtat taacagtggc 562
tggcttgggtg cctggcacct aaaaagcact caataaatgt ttgtttaatg aaaaaaaaaa 622
aaa 625

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<210> 291
<211> 684
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 212..361

<221> sig_peptide
<222> 212..319
<223> Von Heijne matrix
      score 4.09999990463257
      seq HWLFLASLSGIKT/YQ

<221> polyA_signal
<222> 650..655

<221> polyA_site
<222> 673..684

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```

<400> 291
atccccawns cactctctca cagagactgt tcttttcctt ctgagaccct actccagctt 60
gtagttctaa atctgtgatt atgcactgtc tgtcttcctc ttgaggtcag gggccatttc 120
ttttgttctc tgctatgctc aggaccagga tcaaaggagc tcagtaacta tttacaggcg 180
tacatcatat gtggaggaca cttatgctgt g atg gcc cca cac aca gct tcc 232
                               Met Ala Pro His Thr Ala Ser
                               -35                -30
ttt ggg gtc tgt ccc ctg ctg tcc gtt acc cgc gtg gta gcc act gag 280
Phe Gly Val Cys Pro Leu Leu Ser Val Thr Arg Val Val Ala Thr Glu
      -25                -20                -15
cac tgg ctg ttc ctg gct tca ctg tct ggc atc aaa act tat cag tcc 328
His Trp Leu Phe Leu Ala Ser Leu Ser Gly Ile Lys Thr Tyr Gln Ser
      -10                -5                1
tac atc tca gtc ttt tgc aag gtg aca ctt atc tgattaccta attcacacra 381
Tyr Ile Ser Val Phe Cys Lys Val Thr Leu Ile

```

```

5              10
aggtgttaat ggtggtaatg gcataktatt tattacccca ggggaccak aacggtggta 441
tcaaaacata tcattcccca gtggtttaaa actctggtag ctttccargg aatccaaagt 501
ggaatccagt ctcccttagct gawttcacag ggccccgtct gcacaacttg gcttctgtcg 561
gcttccctan ccctgacttc ccaagcetta gtcatacccc tctctccac ccagggtca 621
gcacagtacc tggaacagtc aagccctcaa taaatgttta ctgagtgcac yaaaaaaaaa 681
aaa 684
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<210> 292  
<211> 628  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 75..482

<221> sig\_peptide  
<222> 75..128  
<223> Von Heijne matrix  
score 3.59999990463257  
seq KMLISVAMLGAXA/GV

<221> polyA\_signal  
<222> 595..600

<221> polyA\_site  
<222> 618..627

<400> 292

aa	agtgagacc	gcgcggcaac	agcttgcggc	tgcggggagc	tcccgtaggc	gctccgctgg	60
ctgtgcaggc	ggcc	atg gat tcc ttg	cgg aaa atg	ctg atc tca gtc	gca		110
		Met Asp Ser Leu Arg Lys Met	Leu Ile Ser Val Ala				
		-15	-10				
atg ctg ggc gca	rgg gct ggc	gtg ggc tac	gcg ctc ctc	gtt atc gtg			158
Met Leu Gly Ala	Xaa Ala Gly	Val Gly Tyr	Ala Leu Leu	Val Ile Val			
-5	1	5	10				
acc ccg gga gag	cgg cgg aag	cag gaa atg	cta aag gag	atg cca ctg			206
Thr Pro Gly Glu	Arg Arg Lys	Gln Glu Met	Leu Lys Glu	Met Pro Leu			
	15	20	25				
cag gac cca agg	agc agg gag	gag gcg gcc	agg acc cag	cag cta ttg			254
Gln Asp Pro Arg	Ser Arg Glu	Glu Ala Ala	Arg Thr Gln	Gln Leu Leu			
	30	35	40				
ctg gcc act ctg	cag gag gca	gcg acc acg	cag gag aac	gtg gcc tgg			302
Leu Ala Thr Leu	Gln Glu Ala	Ala Thr Thr	Gln Glu Asn	Val Ala Trp			
	45	50	55				
agg aag aac tgg	atg gtt ggc	ggc gaa ggc	ggc gcc acg	gga kgt cac			350
Arg Lys Asn Trp	Met Val Gly	Gly Gly Glu	Gly Gly Ala	Thr Gly Xaa	His		
60	65	70					
cgt gag acc gga	ctt gcc tcc	gtg ggc gcc	gga cct tgg	ctt ggg cgc			398
Arg Glu Thr Gly	Leu Ala Ser	Val Gly Ala	Gly Pro Trp	Leu Gly Arg			
75	80	85	90				
agg aat ccg agg	cag ctt tct	cct tcg tgg	gcc can cgg	aaa atc cgg			446
Arg Asn Pro Arg	Gln Leu Ser	Pro Ser Trp	Ala Xaa Arg	Lys Ile Arg			
	95	100	105				

```
amc gaa aat wcc atg cca gga ctc tcc ggg gtc ctg tgaactgccg      492
Xaa Glu Asn Xaa Met Pro Gly Leu Ser Gly Val Leu
      110                      115
tcgggtgagc acgtgtcccc caaaccttgg actgactgct ttaaggtccg caaggcgggc      552
cagggccgag acgcgagtcg gatgtggtga actgaaagaa ccaataaaat catgttcctc      612
cammcaaaaa aaaaah                                           628
```

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<210> 293
<211> 813
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 50..631
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<221> sig_peptide
<222> 50..244
<223> Von Heijne matrix
      score 8
      seq LTLIGCLVTGVES/KI
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<221> polyA_signal
<222> 777..782
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```
<221> polyA_site
<222> 801..812
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<400> 293
aaaggaaagga ttactcgagc cttgttagaa tcagacatgg cttcagggg atg cag gac      58
                                         Met Gln Asp
                                         -65
gct ccc ctg agc tgc ctg tca ccg act aag tgg agc agt gtt tct tcc      106
Ala Pro Leu Ser Cys Leu Ser Pro Thr Lys Trp Ser Ser Val Ser Ser
      -60                      -55                      -50
gca gac tca act gag aag tca gcc tct gcg gca ggc acc agg aat ctg      154
Ala Asp Ser Thr Glu Lys Ser Ala Ser Ala Ala Gly Thr Arg Asn Leu
      -45                      -40                      -35
cct ttt cag ttc tgt ctc ccg cag gct ttg agg atg aag gct gcg ggc      202
Pro Phe Gln Phe Cys Leu Arg Gln Ala Leu Arg Met Lys Ala Ala Gly
      -30                      -25                      -20                      -15
att ctg acc ctc att ggc tgc ctg gtc aca ggc gtc gag tcc aaa atc      250
Ile Leu Thr Leu Ile Gly Cys Leu Val Thr Gly Val Glu Ser Lys Ile
      -10                      -5                      1
tac act cgt tgc aaa ctg gca aaa ata ttc tcg agg gct ggc ctg gac      298
Tyr Thr Arg Cys Lys Leu Ala Lys Ile Phe Ser Arg Ala Gly Leu Asp
      5                      10                      15
aat cyg agg ggc ttc agc ctt gga aac tgg atc tgc atg gcg tat tat      346
Asn Xaa Arg Gly Phe Ser Leu Gly Asn Trp Ile Cys Met Ala Tyr Tyr
      20                      25                      30
gag agc ggc tac aac acc aca gcc car acg gtc ctg gat gac ggc agc      394
Glu Ser Gly Tyr Asn Thr Thr Ala Gln Thr Val Leu Asp Asp Gly Ser
      35                      40                      45                      50
atc gac tay ggc atc ttc caa atc aac agc ttc gcg tgg tgc aga cgc      442
Ile Asp Tyr Gly Ile Phe Gln Ile Asn Ser Phe Ala Trp Cys Arg Arg
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	55		60		65	
gga aag ctg aag gag aac aac cac tgc cay gtc gcc tgc tca gcc ttg						490
Gly Lys Leu Lys Glu Asn Asn His Cys His Val Ala Cys Ser Ala Leu						
	70		75		80	
rtc act gat gac ctc aca gat gca att atc tgt gcc arg aaa att gtt						538
Xaa Thr Asp Asp Leu Thr Asp Ala Ile Ile Cys Ala Xaa Lys Ile Val						
	85		90		95	
aaa gag aca caa gga atg aac tat tgg caa ggc tgg aag aaa cay tgt						586
Lys Glu Thr Gln Gly Met Asn Tyr Trp Gln Gly Trp Lys Lys His Cys						
	100		105		110	
gag ggg aga gac ctg tcc gas tgg aaa aaa ggc tgt gag gtt tcc						631
Glu Gly Arg Asp Leu Ser Xaa Trp Lys Lys Gly Cys Glu Val Ser						
	115		120		125	
taaactggaa ctggaccag gatgctttgc ascaacgccc tagggtttgc agtgaatgtc						691
caaatgcctg tgtcatcttg tcccgtttcc tcccaatatt ccttctcaaa cttggagagg						751
gaaaattaag ctatactttt aagaaaataa atattttccat ttaaattgtca amaaaaaaaa						811
ah						813

<210> 294  
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 <221> CDS  
 <222> 154..576  
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 <222> 154..360  
 <223> Von Heijne matrix  
 score 4.80000019073486  
 seq MMVLSLGIILASA/SF  
 <221> polyA\_signal  
 <222> 737..742  
 <221> polyA\_site  
 <222> 763..775

<400> 294	
agtaaaaaaa cactggaata aggaagggct gatgactttc agaagatgaa ggtaagtaga	60
aaccgttgat gggactgaga aaccagagtk aaaacctctt tggagcttct gaggactcag	120
ctggaaccaa cgggcacagt tggcaacacc atc atg aca tca caa cct gtt ccc	174
Met Thr Ser Gln Pro Val Pro	
	-65
aat gag acc atc ata gtg ctc cca tca aat gtc atc aac ttc tcc caa	222
Asn Glu Thr Ile Ile Val Leu Pro Ser Asn Val Ile Asn Phe Ser Gln	
	-60
gca gag aaa ccc gaa ccc acc aac cag ggg cag gat agc ctg aag aaa	270
Ala Glu Lys Pro Glu Pro Thr Asn Gln Gly Gln Asp Ser Leu Lys Lys	
	-45
cat cta cac gca gaa atc aaa gtt att ggg act atc cag atc ttg tgt	318
His Leu His Ala Glu Ile Lys Val Ile Gly Thr Ile Gln Ile Leu Cys	
	-30
ggc atg atg gta ttg agc ttg ggg atc att ttg gca tct gct tcc ttc	366
	-25
	-20
	-15

Gly Met Met Val Leu Ser Leu Gly Ile Ile Leu Ala Ser Ala Ser Phe	
-10 -5 1	
tct cca aat ttt acc caa gtg act tct aca ctg ttg aac tct gct tac	414
Ser Pro Asn Phe Thr Gln Val Thr Ser Thr Leu Leu Asn Ser Ala Tyr	
5 10 15	
cca ttc ata gga ccc ttt ttt gtr akt aaa btt tct gag gag ggc agg	462
Pro Phe Ile Gly Pro Phe Phe Val Xaa Lys Xaa Ser Glu Glu Gly Arg	
20 25 30	
atg ggg caa ara ggg gag gaa rat vcc aat agc tta aac ttc cca sct	510
Met Gly Gln Xaa Gly Glu Glu Xaa Xaa Asn Ser Leu Asn Phe Pro Xaa	
35 40 45 50	
gcc agc ttg cta tkt ttg atc tgc cag gav caa gga ttc aac ggt gaa	558
Ala Ser Leu Leu Xaa Leu Ile Cys Gln Xaa Gln Gly Phe Asn Gly Glu	
55 60 65	
tct tgt tct cct gtc ggg targataaca ggggttgctt ratttttagat	606
Ser Cys Ser Pro Val Gly	
70	
caattttctta tcagactcaa ataaacattt cttttgaaaa tcactcttatt cttcacatta	666
tcactcttgag ctatgatgga aactagtgas ktctctccag gtttaggcga aaaaaaaatc	726
catgaattag gataaagttg ggaaggaaca ttttatacaa aaaaaaaaah cc	778

<210> 295  
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 <222> 154..897  
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 <222> 154..360  
 <223> Von Heijne matrix  
 score 4.80000019073486  
 seq MMVLSLGIILASA/SF

<221> polyA\_signal  
 <222> 1017..1022

<221> polyA\_site  
 <222> 1044..1054

<400> 295	
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aaccgttgat gggactgaga aaccagagtk aaaacctctt tggagcttct gaggactcag	120
ctggaaccaa cgggcacagt tggcaacacc atc atg aca tca caa cct gtt ccc	174
Met Thr Ser Gln Pro Val Pro	
-65	
aat gag acc atc ata gtg ctc cca tca aat gtc atc aac ttc tcc caa	222
Asn Glu Thr Ile Ile Val Leu Pro Ser Asn Val Ile Asn Phe Ser Gln	
-60 -55 -50	
gca gag aaa ccc gaa ccc acc aac cag ggg cag gat agc ctg aag aaa	270
Ala Glu Lys Pro Glu Pro Thr Asn Gln Gly Gln Asp Ser Leu Lys Lys	
-45 -40 -35	
cat cta cac gca gar rtc aaa gtt att ggg act atc cag atc ttg tgt	318



His	Leu	His	Ala	Glu	Xaa	Lys	Val	Ile	Gly	Thr	Ile	Gln	Ile	Leu	Cys	
-30					-25					-20					-15	
ggc	atg	atg	gta	ttg	agc	ttg	ggg	atc	att	ttg	gca	tct	gct	tcc	ttc	366
Gly	Met	Met	Val	Leu	Ser	Leu	Gly	Ile	Ile	Leu	Ala	Ser	Ala	Ser	Phe	
			-10					-5						1		
tct	cca	aat	ttt	acc	caa	gtg	act	tct	aca	ctg	ttg	aac	tct	gct	tac	414
Ser	Pro	Asn	Phe	Thr	Gln	Val	Thr	Ser	Thr	Leu	Leu	Asn	Ser	Ala	Tyr	
	5					10				15						
cca	ttc	ata	gga	ccc	ttt	ttt	ttt	atc	atc	tct	ggc	tct	cta	tca	atc	462
Pro	Phe	Ile	Gly	Pro	Phe	Phe	Phe	Ile	Ile	Ser	Gly	Ser	Leu	Ser	Ile	
	20				25					30						
gcc	aca	aaa	aaa	agg	tta	acc	aac	ctt	ttg	gtg	cat	acc	acc	ctg	gtt	510
Ala	Thr	Lys	Lys	Arg	Leu	Thr	Asn	Leu	Leu	Val	His	Thr	Thr	Leu	Val	
35				40				45						50		
gga	agc	att	ctg	agt	gct	ctg	tct	gcc	ctg	gtg	ggg	ttc	att	ayc	ctg	558
Gly	Ser	Ile	Leu	Ser	Ala	Leu	Ser	Ala	Leu	Val	Gly	Phe	Ile	Xaa	Leu	
			55					60						65		
tct	gtc	aaa	cag	gcc	acc	tta	aat	cct	gcc	tca	ctg	cak	tgt	gag	ttg	606
Ser	Val	Lys	Gln	Ala	Thr	Leu	Asn	Pro	Ala	Ser	Leu	Xaa	Cys	Glu	Leu	
			70				75						80			
gmc	aaa	aat	aat	ata	cca	aca	ara	akt	tat	gtt	yct	tac	ttt	tat	cat	654
Xaa	Lys	Asn	Asn	Ile	Pro	Thr	Xaa	Xaa	Tyr	Val	Xaa	Tyr	Phe	Tyr	His	
		85				90						95				
gat	tca	ctt	tat	acc	acg	gac	kgc	tat	aca	gcc	aaa	gcc	akt	ctg	gct	702
Asp	Ser	Leu	Tyr	Thr	Thr	Asp	Xaa	Tyr	Thr	Ala	Lys	Ala	Xaa	Leu	Ala	
	100					105				110						
gga	act	ctc	tct	ctg	atg	ctg	att	tgc	act	ctg	ctg	gaa	ttc	tgc	cwa	750
Gly	Thr	Leu	Ser	Leu	Met	Leu	Ile	Cys	Thr	Leu	Leu	Glu	Phe	Cys	Xaa	
115				120				125						130		
act	gtg	ctc	act	gct	gtg	ctg	cgg	tgg	aaa	cag	gct	tac	tct	gac	ttc	798
Xaa	Val	Leu	Thr	Ala	Val	Leu	Arg	Trp	Lys	Gln	Ala	Tyr	Ser	Asp	Phe	
			135				140							145		
cct	ggg	agt	gta	ctt	ttc	ctg	cct	cam	agt	tac	att	ggw	aat	tct	ggm	846
Pro	Gly	Ser	Val	Leu	Phe	Leu	Pro	Xaa	Ser	Tyr	Ile	Gly	Asn	Ser	Gly	
		150					155						160			
atg	tcc	tca	aaa	atg	acy	cat	gac	tgt	gga	tat	gaa	gaa	cta	ttg	act	894
Met	Ser	Ser	Lys	Met	Thr	His	Asp	Cys	Gly	Tyr	Glu	Glu	Leu	Leu	Thr	
	165					170					175					
tct	taagaaaaaa	gggagaaata	ttaatcagaa	agttgattct	tatgataata											947
Ser																
tggaagtt	aaccattata	gaaaagcaaa	gcttgagttt	cctaaatgta	agctttttaa											1007
gtaatgaaca	ttaaaaaaa	ccattatttc	actgtcaaaa	aaaaaaamcc	nkt											1060

<210> 296  
 <211> 444  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 146..292  
 <221> sig\_peptide  
 <222> 146..253  
 <223> Von Heijne matrix

score 5.5  
seq FTSMCILFHCLLS/FQ

<221> polyA\_signal  
<222> 395..400

<221> polyA\_site  
<222> 433..444

<400> 296  
aacttgggac aagaratcaa actttaaga tgggtctaaag cccctcttaa aggtctgact 60  
gtgtcgggacc tctagagcta atctcactag atgtgagcca ttgtttatat tctagccatc 120  
ctttcattttc attctagaag accccc atg caa gtt ccc cac cta agg gtc tgg 172  
Met Gln Val Pro His Leu Arg Val Trp  
-35 -30  
aca cag gtg awa gat acc ttc att ggt tat aga aat ttg gga ttt aca 220  
Thr Gln Val Xaa Asp Thr Phe Ile Gly Tyr Arg Asn Leu Gly Phe Thr  
-25 -20 -15  
agt atg tgc ata ttg ttc cac tgt ctt ctt agc ttt cag gtt ttc aaa 268  
Ser Met Cys Ile Leu Phe His Cys Leu Leu Ser Phe Gln Val Phe Lys  
-10 -5 1 5  
aag aaa aga aaa ctt ara ctt ttc tgatgttctt ttttacgtaa ataaccattt 322  
Lys Lys Arg Lys Leu Xaa Leu Phe  
10  
tattgtttgtt ttgttttttc tgccttcaaa ctactccac aggccaaata tavctggctg 382  
cttctttctg taaataaagt tttattgggc cacagccatg gccatctttt aaaaaaaaaa 442  
aa 444

<210> 297  
<211> 754  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 126..383

<221> sig\_peptide  
<222> 126..167  
<223> Von Heijne matrix  
score 7.5  
seq VALNLILVPCCAA/WC

<221> polyA\_signal  
<222> 726..731

<221> polyA\_site  
<222> 743..754

<400> 297  
aattgtatgt tacgatgttg tattgatttt taagaaagta attkratttg taaaacttct 60  
gctcgttttac actgcacatt gaatacaggt aactaattgg wwgagaggg gaggtcactc 120  
ttttg atg gtg gcc ctg aac ctc att ctg gtt ccc tgc tgc gct gct tgg 170  
Met Val Ala Leu Asn Leu Ile Leu Val Pro Cys Cys Ala Ala Trp  
-10 -5 1

tgt gac cca cgg agg atc cac tcc cag gat gac gtg ctc cgt agc tct	218
Cys Asp Pro Arg Arg Ile His Ser Gln Asp Asp Val Leu Arg Ser Ser	
5 10 15	
gct gct gat act ggg tct gcg atg cag cgg cgt gag gcc tgg gct ggt	266
Ala Ala Asp Thr Gly Ser Ala Met Gln Arg Arg Glu Ala Trp Ala Gly	
20 25 30	
tgg aga agg tca caa ccc ttc tct gtt ggt ctg cct tct gct gaa aga	314
Trp Arg Arg Ser Gln Pro Phe Ser Val Gly Leu Pro Ser Ala Glu Arg	
35 40 45	
ctc gag aac caa cca ggg aag ctg tcc tgg agg tcc ctg gtc gga gag	362
Leu Glu Asn Gln Pro Gly Lys Leu Ser Trp Arg Ser Leu Val Gly Glu	
50 55 60 65	
gga cat aga atc tgt gac ctc tgacrrctgt gaasccaccc tgggctacar	413
Gly His Arg Ile Cys Asp Leu	
70	
aaaccacagt cttcccagca attattacaa ttcttgaatt ccttggggat tttttactgc	473
cctttcaaag cacttaaktg tkrratctaa cgtkttccag tgtctgtctg aggtgactta	533
aaaaatcaga acaaaacttc tattatccag agtcatggga gagtacaccc tttccaggaa	593
taatgttttg ggaaacactg aaatgaaatc ttcccagtat tataaattgt gtatttaaaa	653
aaaagaaact tttctgaatg cctacctggc ggtgtatacc aggcagtgtg ccagtttaaa	713
aagatgaaaa agaataaaaa cttttgagga aaaaaaaaa a	754

<210> 298  
 <211> 629  
 <212> DNA  
 <213> Homo sapiens  
 <220>  
 <221> CDS  
 <222> 66..497  
 <221> sig\_peptide  
 <222> 66..239  
 <223> Von Heijne matrix  
 score 5.40000009536743  
 seq QLLDSVLWLGLG/LT

<221> polyA\_signal  
 <222> 594..599

<221> polyA\_site  
 <222> 618..629

<400> 298	
aactcccaga atgetgacca aagtgggagg agcactaggt cttcccgctca cctccacctc	60
tctcc atg acc cgg ctc tgc tta ccc aga ccc gaa gca cgt gag gat ccg	110
Met Thr Arg Leu Cys Leu Pro Arg Pro Glu Ala Arg Glu Asp Pro	
-55 -50 -45	
atc cca gtt cct cca agg ggc ctg ggt gct ggg gag ggg tca ggt agt	158
Ile Pro Val Pro Pro Arg Gly Leu Gly Ala Gly Glu Gly Ser Gly Ser	
-40 -35 -30	
cca gtg cgt cca cct gta tcc acc tgg ggc cct agc tgg gcc cag ctc	206
Pro Val Arg Pro Pro Val Ser Thr Trp Gly Pro Ser Trp Ala Gln Leu	
-25 -20 -15	
ctg gac agt gtc cta tgg ctg ggg gca cta gga ctg aca atc cag gca	254

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Leu Asp Ser Val Leu Trp Leu Gly Ala Leu Gly Leu Thr Ile Gln Ala
-10 -5 1 5
gtc ttt tcc acc act ggc cca gcc ctg ctg ctg ctt ctg gtc agc ttc 302
Val Phe Ser Thr Thr Gly Pro Ala Leu Leu Leu Leu Val Ser Phe
10 15 20
ctc acc ttt gac ctg ctc cat agg ccc gca gtc aca ctc tgc cac agc 350
Leu Thr Phe Asp Leu Leu His Arg Pro Ala Val Thr Leu Cys His Ser
25 30 35
gca aac ttc tca cca ggg gcc aga gtc agg ggg ccg gtg aag gtc ctg 398
Ala Asn Phe Ser Pro Gly Ala Arg Val Arg Gly Pro Val Lys Val Leu
40 45 50
gac agc agg agg ctc tac tcc tgc aaa tgg gta cag tct cag gac aac 446
Asp Ser Arg Arg Leu Tyr Ser Cys Lys Trp Val Gln Ser Gln Asp Asn
55 60 65
tta gcc tcc agg aag cac tgc tgc tgc tgc tca tgg ggc tgg gcc cgc 494
Leu Ala Ser Arg Lys His Cys Cys Cys Cys Ser Trp Gly Trp Ala Arg
70 75 80 85
tcc tgaaaacctg tggcatgccc ttgwaccctg cttggcctgg ctttctgcct 547
Ser
ccatccttgg gcctgakanc ccctccccac aactcagtgt ccttcaaata tacaatgacc 607
acccttcttc aaaaaaaaaa aa 629

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210> 299
211> 765
212> DNA
213> Homo sapiens
220>
221> CDS
222> 49..411
221> sig_peptide
222> 49..96
223> Von Heijne matrix
score 10.1000003814697
seq LVLTLCTLPLAVA/SA

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<221> polyA_signal
<222> 732..737

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<221> polyA_site
<222> 750..763

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<400> 299
aaagatccct gcagcccggc aggagagaag gctgagcctt ctggcgtc atg gag agg 57
Met Glu Arg
-15
ctc gtc cta acc ctg tgc acc ctc ccg ctg gct gtg gcg tct gct ggc 105
Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala Ser Ala Gly
-10 -5 1
tgc gcc acg acg cca gct cgc aac ctg agc tgc tac cag tgc ttc aag 153
Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln Cys Phe Lys
5 10 15
gtc agc agc tgg acg gag tgc ccg ccc acc tgg tgc agc ccg ctg gac 201
Val Ser Ser Trp Thr Glu Cys Pro Pro Thr Trp Cys Ser Pro Leu Asp

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20	25	30	35	
caa gtc tgc atc tcc aac gag gtg gtc gtc tct ttt agt gag tcy ccc				249
Gln Val Cys Ile Ser Asn Glu Val Val Val Ser Phe Ser Glu Ser Pro				
	40	45	50	
ccg ggc aga ggg cas gtg cca bgt gcc ggg gaa kgg ccg gtg ccc ccg				297
Pro Gly Arg Gly Xaa Val Pro Xaa Ala Gly Glu Xaa Pro Val Pro Pro				
	55	60	65	
cct ctc wkc gac tta bct atg act cct cgg ckc ycc agg gcc tgg ggc				345
Pro Leu Xaa Asp Leu Xaa Met Thr Pro Arg Xaa Xaa Arg Ala Trp Gly				
	70	75	80	
ckk gtg ggt ccd aaa gtg cct cct gct gtc tct ccc gcg ctg ggc tcg				393
Pro Val Gly Pro Lys Val Pro Pro Ala Val Ser Pro Ala Leu Gly Ser				
	85	90	95	
ggc gag cat ccs rva btg tgaatkkkga cttttttctc ckccatttga				441
Gly Glu His Pro Xaa Xaa				
100	105			
agtgtcacta ggaactgtca gcaggacaaa ggctctgatg tcaactgaatt tacaaaraca				501
gcaggaacrs ackggtgggg atgggcagct gtterargcr atggggtkac tgcccttcct				561
ggcacagcac artacacctg ccatacaacc carcatcagg cakgctgcac tggaatcgat				621
acagtgtatg acaatgtcat atagtataac acaacataat gaatataacg tgtatattgc				681
aacttaatat aatacgatgt aatataatgc tacataatac aacataatat aataaaatag				741
aatgcaacac aaaaaaaaaa aacc				765

<210> 300

<211> 623

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 49..534

<221> sig\_peptide

<222> 49..96

<223> Von Heijne matrix

score 10.1000003814697

seq LVLTLCTLPLAVA/SA

<221> polyA\_signal

<222> 593..598

<221> polyA\_site

<222> 612..623

<400> 300

aaagatccct gcagcccggc aggagagaag gctgagcctt ctggcgtc atg gag agg	57
Met Glu Arg	
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ctc gtc cta acc ctg tgc acc ctc ccg ctg gct gtg gcg tct gct ggc	105
Leu Val Leu Thr Leu Cys Thr Leu Pro Leu Ala Val Ala Ser Ala Gly	
-10	
-5	
1	
tgc gcc acg acg cca gct cgc aac ctg agc tgc tac cag tgc ttc aag	153
Cys Ala Thr Thr Pro Ala Arg Asn Leu Ser Cys Tyr Gln Cys Phe Lys	
5	
10	
15	
gtc agc agc tgg acg gag tgc ccg ccc acc tgg tgc agc ccg ctg gac	201

Val	Ser	Ser	Trp	Thr	Glu	Cys	Pro	Pro	Thr	Trp	Cys	Ser	Pro	Leu	Asp	
20					25					30					35	
caa	gtc	tgc	atc	tcc	aac	gag	gtg	gtc	gtc	tct	ttt	aaa	tgg	agt	gta	249
Gln	Val	Cys	Ile	Ser	Asn	Glu	Val	Val	Val	Ser	Phe	Lys	Trp	Ser	Val	
			40					45					50			
cgc	gtc	ctg	ctc	agc	aaa	cgc	tgt	gct	ccc	aga	tgt	ccc	aac	gac	aac	297
Arg	Val	Leu	Leu	Ser	Lys	Arg	Cys	Ala	Pro	Arg	Cys	Pro	Asn	Asp	Asn	
		55				60						65				
atg	aak	ttc	gaa	tgg	tgc	ccg	gcc	ccc	atg	gtg	caa	ggc	gtg	atc	acc	345
Met	Xaa	Phe	Glu	Trp	Ser	Pro	Ala	Pro	Met	Val	Gln	Gly	Val	Ile	Thr	
	70					75					80					
agg	cgc	tgc	tgt	tcc	tgg	gct	ctc	tgc	aac	agg	gca	ctg	acc	cca	cag	393
Arg	Arg	Cys	Cys	Ser	Trp	Ala	Leu	Cys	Asn	Arg	Ala	Leu	Thr	Pro	Gln	
	85				90					95						
gag	ggg	cgc	tgg	gcc	ctg	cra	ggg	ggg	ctc	ctg	ctc	cag	gac	cct	tcg	441
Glu	Gly	Arg	Trp	Ala	Leu	Xaa	Gly	Gly	Leu	Leu	Leu	Gln	Asp	Pro	Ser	
100			105					110						115		
agg	ggc	ara	aaa	acc	tgg	gtg	cgg	cca	cag	ctg	ggg	ctc	cca	ctc	tgc	489
Arg	Gly	Xaa	Lys	Thr	Trp	Val	Arg	Pro	Gln	Leu	Gly	Leu	Pro	Leu	Cys	
		120				125						130				
ctt	ccc	awt	tcc	aac	ccc	ctc	tgc	cca	rgg	gaa	acc	cag	gaa	gga		534
Leu	Pro	Xaa	Ser	Asn	Pro	Leu	Cys	Pro	Xaa	Glu	Thr	Gln	Glu	Gly		
		135				140						145				
taacactgtg	ggtgccccca	cctgtgcatt	gggaccacra	cttcaccctc	ttggaracaa											594
taaaactctca	tgcccccaaaa	aaaaaaaaa														623

<210> 301

<211> 571

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 86..415

<221> sig\_peptide

<222> 86..145

<223> Von Heijne matrix

score 9.80000019073486

seq FTIGLTLLLGXQA/MP

<221> polyA\_signal

<222> 540..545

<221> polyA\_site

<222> 560..571

<400> 301

aaaaactcac ccagtgagtg tgagcattta agaagcatcc tctgccaaga ccaaaaggaa 60

agaagaaaaa bggccaaaag ccaaa atg ara ctg atg gta ctt gtt ttc acc 112

Met Xaa Leu Met Val Leu Val Phe Thr

-20 -15

att ggg cta act ttg ctg cta gga rtt caa gcc atg cct gca aat cgc 160

Ile Gly Leu Thr Leu Leu Leu Gly Xaa Gln Ala Met Pro Ala Asn Arg

-10

-5

1

5

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ctc tct tgc tac aga aag ata cta aaa gat cac aac tgt cac aac ctt      208
Leu Ser Cys Tyr Arg Lys Ile Leu Lys Asp His Asn Cys His Asn Leu
      10                      15                      20
ccg gaa gga gta gct gac ctg aca cag att gat gtc aat gtc cag gat      256
Pro Glu Gly Val Ala Asp Leu Thr Gln Ile Asp Val Asn Val Gln Asp
      25                      30                      35
cat ttc tgg gat ggg aag gga tgt gag atg atc tgt tac tgc aac ttc      304
His Phe Trp Asp Gly Lys Gly Cys Glu Met Ile Cys Tyr Cys Asn Phe
      40                      45                      50
aag cga att gct ctg ctg ccc aaa aga cgt ttt ctt tgg acc aaa gat      352
Lys Arg Ile Ala Leu Leu Pro Lys Arg Arg Phe Leu Trp Thr Lys Asp
      55                      60                      65
ctc ttt cgt gat tcc ttg caa caa tca atg aga atc ttc atg tat tct      400
Leu Phe Arg Asp Ser Leu Gln Gln Ser Met Arg Ile Phe Met Tyr Ser
      70                      75                      80                      85
ggc gaa cac cat tcc tgatttccca caaactgcac tacatcagta taactgcatt      455
Gly Glu His His Ser
      90
tctagtttct atatagtgc atagagcata gattctataa attcttactt gtctaagaaa      515
gtaaatctgt gttaaacaag tagtaataaaa agttaattca atccaaaaaa aaaaaa      571

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```

<210> 302
<211> 612
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 56..268
<221> sig_peptide
<222> 56..100
<223> Von Heijne matrix
      score 4.59999990463257
      seq LLTHNLLSSHVRG/VG
<221> polyA_signal
<222> 584..589
<221> polyA_site
<222> 601..612

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<400> 302
ctaatacgaag aggggggattt tccgggttccg gcctggcgag agtttgtgcg gcgac atg      58
                                     Met
                                     -15
aaa ctg ctt acc cac aat ctg ctg agc tgc cat gtg cgg ggg gtg ggg      106
Lys Leu Leu Thr His Asn Leu Leu Ser Ser His Val Arg Gly Val Gly
      -10                      -5                      1
tcc cgt ggc ttc ccc ctg cgc ctc cag gcc acc gag gtc cgt atc tgc      154
Ser Arg Gly Phe Pro Leu Arg Leu Gln Ala Thr Glu Val Arg Ile Cys
      5                      10                      15
cct gtg gaa ttc aac ccc aac ttc gtg gcg cgt atg ata cct aaa gtg      202
Pro Val Glu Phe Asn Pro Asn Phe Val Ala Arg Met Ile Pro Lys Val
      20                      25                      30

```

```
gag tgg tcg gcg ttc ctg gag gcg rmc gat aac ttg cgt ctg atc cag      250
Glu Trp Ser Ala Phe Leu Glu Ala Xaa Asp Asn Leu Arg Leu Ile Gln
35              40              45              50
gtg ccg aga agg gcc ggt tgagggatat gaggagaatg aggagtttct      298
Val Pro Arg Arg Ala Gly
55
gaggaccatg caccacctgc tgctggaggt ggamstgaka gagggcaccc tgcagtgccc      358
ggaatctgga cgtatgttcc ccatcagccg cgggatcccc aacatgctgc tgagtgaaga      418
ggaaactgag agttgattgt gccaggcgcc agtttttctt gttatgactg tgtatttttg      478
ttgatctata ccctgtttcc gaattctgcc gtgtgtatcc ccaacccttg acccaatgac      538
accaaaacaca gtgtttttga gctcgggtatt atatattttt ttctcattaa aggtttaaaa      598
ccaaaaaaaaaaaaa      612
```

```
<210> 303
<211> 539
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 32..328
<221> sig_peptide
<222> 32..103
<223> Von Heijne matrix
      score 4.59999990463257
      seq FFIFCSLNTLLLG/GV
<221> polyA_signal
<222> 508..513
<221> polyA_site
<222> 528..539
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```
K400> 303
aacaaactatc ctgcctgctg cttgctgcac c atg aag tct gcc aag ctg gga      52
Met Lys Ser Ala Lys Leu Gly
-20
ttt ctt cta aga ttc ttc atc ttc tgc tca ttg aat acc ctg tta ttg      100
Phe Leu Leu Arg Phe Phe Ile Phe Cys Ser Leu Asn Thr Leu Leu Leu
-15 -10 -5
ggt ggt gtt aat aaa att gcg gag aag ata tgt gga gac ctc aaa gat      148
Gly Gly Val Asn Lys Ile Ala Glu Lys Ile Cys Gly Asp Leu Lys Asp
1 5 10 15
ccc tgc aaa ttg gac atg aat ttt gga agc tgc tat gaa gtt cac ttt      196
Pro Cys Lys Leu Asp Met Asn Phe Gly Ser Cys Tyr Glu Val His Phe
20 25 30
aga tat ttc tac aac aga acc tcc aaa aga tgt gaa act ttt gtc ttc      244
Arg Tyr Phe Tyr Asn Arg Thr Ser Lys Arg Cys Glu Thr Phe Val Phe
35 40 45
tcc agc tgt aat ggc aac ctt aac aac ttc aag ctt aaa ata gaa cgt      292
Ser Ser Cys Asn Gly Asn Leu Asn Asn Phe Lys Leu Lys Ile Glu Arg
50 55 60
gaa gta kcc tgt gtt gca aaa tac aaa cca ccg agg tgagaggatg      338
Glu Val Xaa Cys Val Ala Lys Tyr Lys Pro Pro Arg
```



65	70	75	
tgaactcatg	aagttgtctg	ctgcaccatc	cgaaataaag acacaagaaa attcaractg 398
attdtgaaat	ctttgttwtg	tttccmymak	ggcgwktaag cttccatattg tttgctattd 458
tcctgaccct	agttttgtct	ttcctggaaa	ttactgtat gacattasa atgaaagagt 518
ctttctgtca	aaaaaaaaa	a	539

<210> 304  
<211> 964  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 21..527

<221> sig\_peptide  
<222> 21..95  
<223> Von Heijne matrix  
score 8.5  
seq LKVLPLAPAAA/QD

<221> polyA\_signal  
<222> 921..926

<221> polyA\_site  
<222> 953..963

<400> 304	
agggcggtatc ttctccggcc atg agg aag cca gcc gct ggc ttc ctt ccc tca	53
Met Arg Lys Pro Ala Ala Gly Phe Leu Pro Ser	
-25 -20 -15	
ctc ctg aag gtg ctg ctc ctg cct ctg gca cct gcc gca gcc cag gat	101
Leu Leu Lys Val Leu Leu Pro Leu Ala Pro Ala Ala Gln Asp	
-10 -5 1	
acg act cag gcc tcc act cca ggc agc cct ctc tct cct acc gaa tac	149
Ser Thr Gln Ala Ser Thr Pro Gly Ser Pro Leu Ser Pro Thr Glu Tyr	
5 10 15	
caa cgc ttc ttc gca ctg ctg act cca acc tgg aag gca gar act acc	197
Gln Arg Phe Phe Ala Leu Leu Thr Pro Thr Trp Lys Ala Glu Thr Thr	
20 25 30	
tgc cgt ctc cgt gca acc cac ggc tgc cgg aat ccc aca ctc gtc cag	245
Cys Arg Leu Arg Ala Thr His Gly Cys Arg Asn Pro Thr Leu Val Gln	
35 40 45 50	
ctg gac caa tat gaa aac cac ggc tta gtg ccc gat ggt gct gtc tgc	293
Leu Asp Gln Tyr Glu Asn His Gly Leu Val Pro Asp Gly Ala Val Cys	
55 60 65	
tcc aac ctc cct tat gcc tcc tgg ttt gag tct ttc tgc cag ttc act	341
Ser Asn Leu Pro Tyr Ala Ser Trp Phe Glu Ser Phe Cys Gln Phe Thr	
70 75 80	
cac tac cgt tgc tcc aac cac gtc tac tat gcc aag aga gtc ctg tgt	389
His Tyr Arg Cys Ser Asn His Val Tyr Tyr Ala Lys Arg Val Leu Cys	
85 90 95	
tcc cag cca gtc tct att ctc tcw cct aac act ctc aag gag ata gaa	437
Ser Gln Pro Val Ser Ile Leu Ser Pro Asn Thr Leu Lys Glu Ile Glu	
100 105 110	

```

sct tca gct gaa gtc tca ccc acc aca gat gac ctc ccc cat ctc acc 485
Xaa Ser Ala Glu Val Ser Pro Thr Thr Asp Asp Leu Pro His Leu Thr
115          120          125          130
cca ctt cac agt gac aga acg cca gac ctt cca gcc ctg gcc 527
Pro Leu His Ser Asp Arg Thr Pro Asp Leu Pro Ala Leu Ala
135          140
tgagaggctc agcaacaacg tggaagagct cctacaatcc tccttggtccc tgggaggcca 587
ggagcaagcg ccagagcaca agcaggagca aggagtggag cacaggcagg agccgacaca 647
agaacacaag caggaagagg ggcagaaaca ggaagagcaa gaagaggaac aggaagagga 707
gggaaagcag gaagaaggac aggggactaa ggagggacgg gaggctgtgt ctcagctgca 767
gacagactca gagcccaagt ttactctga atctctatct tctaaccctt cctcttttgc 827
tccccgggta cganaagtag agtctactcc tatgataatg gagaacatcc aggagctcat 887
tcgatcagcc caggaaatag atgaaatgaa tgaaatatat gatgagaact cctactggag 947
aaacccaaaa aaaaaak 964

```

```

<210> 305
<211> 684
<212> DNA
<213> Homo sapiens

```

```

<220>
<221> CDS
<222> 147..647
<221> sig_peptide
<222> 147..374
<223> Von Heijne matrix
      score 3.5
      seq LASASELPLGSRP/AP

```

```

<221> polyA_site
<222> 668..681

```

```

<400> 305
aacttcctgt gagcccggcg gtgacaacgg caacatggcc cgtgaacgga gctgaagtcg 60
acgacttctc ctrgrarmcc ccgactgagg cggagacgaa ggtgctgcag gcgcgacggg 120
agcggcaaga tcgcatctcc cggtctc atg ggc gac tat ctg ctg cgc ggt tac 173
          Met Gly Asp Tyr Leu Leu Arg Gly Tyr
          -75          -70
cgc atg ctg ggc gag acg tgt gcg gac tgc ggg acg atc ctc ctc caa 221
Arg Met Leu Gly Glu Thr Cys Ala Asp Cys Gly Thr Ile Leu Leu Gln
          -65          -60          -55
gac aaa cag cgg aaa atc tac tgc gtg gct tgt cag gaa ctc gac tca 269
Asp Lys Gln Arg Lys Ile Tyr Cys Val Ala Cys Gln Glu Leu Asp Ser
          -50          -45          -40
gac gtg gat aaa gat aat ccc gct ctg aat gcc cag gct gcc ctc tcc 317
Asp Val Asp Lys Asp Asn Pro Ala Leu Asn Ala Gln Ala Ala Leu Ser
          -35          -30          -25          -20
caa gct cgg gag cac cag ctg gcc tca gcc tca gag ctc ccc ctg ggc 365
Gln Ala Arg Glu His Gln Leu Ala Ser Ala Ser Glu Leu Pro Leu Gly
          -15          -10          -5
tct cga cct gcg ccc caa ccc cca gta cct cgt ccg gag cac tgt gag 413
Ser Arg Pro Ala Pro Gln Pro Pro Val Pro Arg Pro Glu His Cys Glu
          1          5          10
gga gct gca gca gga ctc aag gca gcc cag ggg cca cct gct cct gct 461

```

[illegible]

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<210> 306
<211> 693
<212> DNA
<213> Homo sapiens
```

$\langle 220 \rangle$   
 $\langle 221 \rangle$  CDS  
 $\langle 222 \rangle$  262..471

```
<221> sig_peptide
<222> 262..306
<223> Von Heijne matrix
      score 3.5
      seq LCFLLPHHRLQEA/RQ
```

```
<221> polyA_signal
<222> 663..668
```

```
<221> polyA_site
<222> 682..693
```

[illegible]

Gly Asn Xaa Xaa Xaa Xaa Xaa Leu Ser Lys Arg Asp  
45 50 55  
ctgacccttt tgatttccaa vctcascgtt ttggtgtaag gcggccaaar aaggatgcgg 541  
ascccagcac tgtgaagcct acaaaaacat tgatgcgctg gcttggggat ttgaatttga 601  
acatctttca cactaagttc agactcatga aaccaatctt cagatgctct gtaaaccaca 661  
taataaagag tttggaaatt aaaaaaaaaa aa 693

<210> 307  
<211> 1656  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 74..1216

<221> sig\_peptide  
<222> 74..172  
<223> Von Heijne matrix  
score 5.80000019073486  
seq XLCLGMALCPRQA/TR

<221> polyA\_signal  
<222> 1627..1632

<221> polyA\_site  
<222> 1640..1652

<400> 307  
atctcttggc gtctcaacgt tcggatcagc agcttttttc cattctctct ctccacttct 60  
tcagtgaagc gcc atg agt tgg act gtg cct gtt gtg cgg gcc agc cag 109  
Met Ser Trp Thr Val Pro Val Val Arg Ala Ser Gln  
-30 -25  
aga gtg agc tcg gtg gga gcg aat ktc cta tgc ctg ggg atg gcc ctg 157  
Arg Val Ser Ser Val Gly Ala Asn Xaa Leu Cys Leu Gly Met Ala Leu  
-20 -15 -10  
tgt ccg cgt caa gca acg cgc atc ccg ctc aac ggc acc tgg ctc ttc 205  
Cys Pro Arg Gln Ala Thr Arg Ile Pro Leu Asn Gly Thr Trp Leu Phe  
-5 1 5 10  
acc ccc gtg agc aag atg gcg act gtg aar agt gag ctt att gag cgt 253  
Thr Pro Val Ser Lys Met Ala Thr Val Lys Ser Glu Leu Ile Glu Arg  
15 20 25  
ttc act tcc gar aag ccc gtt cat cac agt aag gtc tcc atc ata gga 301  
Phe Thr Ser Glu Lys Pro Val His His Ser Lys Val Ser Ile Ile Gly  
30 35 40  
act gga tcg gtg ggc atg gcc tgc gct atc agc atc tta tta aaa ggc 349  
Thr Gly Ser Val Gly Met Ala Cys Ala Ile Ser Ile Leu Leu Lys Gly  
45 50 55  
ttg agt gat gaa ctt gcc ctt gtg gat ctt gat gaa rac aaa ctg aag 397  
Leu Ser Asp Glu Leu Ala Leu Val Asp Leu Asp Glu Xaa Lys Leu Lys  
60 65 70 75  
ggg gag acr atg gat ctt caa cat ggc agc cct ttc acg aaa atg cca 445  
Gly Glu Thr Met Asp Leu Gln His Gly Ser Pro Phe Thr Lys Met Pro  
80 85 90  
aat att gtt tgt agc aaa rat tac ttt gtc aca gca aac tcc aac cta 493

Asn Ile Val Cys Ser Lys Xaa Tyr Phe Val Thr Ala Asn Ser Asn Leu	
95 100 105	
gtg att atc aca gca ggt gca cgc caa raa aag gga gaa acg cgc ctt	541
Val Ile Ile Thr Ala Gly Ala Arg Gln Xaa Lys Gly Glu Thr Arg Leu	
110 115 120	
aat tta stc cag cga aat gtg gcc atc ttc aag tta atg att tcc agt	589
Asn Leu Xaa Gln Arg Asn Val Ala Ile Phe Lys Leu Met Ile Ser Ser	
125 130 135	
att gtc cag tac agc ccc cac tgc aaa ctg att att gtt tcc aat cca	637
Ile Val Gln Tyr Ser Pro His Cys Lys Leu Ile Ile Val Ser Asn Pro	
140 145 150 155	
gtg gat atc tta act tat gta gct tgg aag ttg agt gca ttt ccc aaa	685
Val Asp Ile Leu Thr Tyr Val Ala Trp Lys Leu Ser Ala Phe Pro Lys	
160 165 170	
aac cgt att att gga agc ggc tgt aat ctg ata mhg gct cgt ttt cgt	733
Asn Arg Ile Ile Gly Ser Gly Cys Asn Leu Ile Xaa Ala Arg Phe Arg	
175 180 185	
ttc ttg att gga caa aag ctt ggt atc cat tct gaa agc tgc cat gga	781
Phe Leu Ile Gly Gln Lys Leu Gly Ile His Ser Glu Ser Cys His Gly	
190 195 200	
tgg atc ctc gga gag cat gga gac tca agt gtt cct gtg tgg agt gga	829
Trp Ile Leu Gly Glu His Gly Asp Ser Ser Val Pro Val Trp Ser Gly	
205 210 215	
gtg aac ata gct ggt gtc cct ttg aag gat ctg aac tct gat ata gga	877
Val Asn Ile Ala Gly Val Pro Leu Lys Asp Leu Asn Ser Asp Ile Gly	
220 225 230 235	
act gat aaa gat cct gag caa tgg aaa aat gtc cac aaa gaa gtg act	925
Thr Asp Lys Asp Pro Glu Gln Trp Lys Asn Val His Lys Glu Val Thr	
240 245 250	
gca act gcc tat gag att att aaa atg aaa ggt tat act tct tgg gcc	973
Ala Thr Ala Tyr Glu Ile Ile Lys Met Lys Gly Tyr Thr Ser Trp Ala	
255 260 265	
att ggc cta tct gtg gcc gat tta aca gaa agt att ttg aag aat ctt	1021
Ile Gly Leu Ser Val Ala Asp Leu Thr Glu Ser Ile Leu Lys Asn Leu	
270 275 280	
agg aga ata cat cca gtt tcc acc ata act aag ggc ctc tat gga ata	1069
Arg Arg Ile His Pro Val Ser Thr Ile Thr Lys Gly Leu Tyr Gly Ile	
285 290 295	
rat gaa gaa gta ttc ctc agt att cct tgt atc ctg gga gag aac ggt	1117
Xaa Glu Glu Val Phe Leu Ser Ile Pro Cys Ile Leu Gly Glu Asn Gly	
300 305 310 315	
att acc aac ctt ata aag ata aag ctg acc cct gaa gaa gag gcc cat	1165
Ile Thr Asn Leu Ile Lys Ile Lys Leu Thr Pro Glu Glu Glu Ala His	
320 325 330	
ctg aaa aaa agt gca aaa aca ctc tgg gaa att cag aat aag ctt aag	1213
Leu Lys Lys Ser Ala Lys Thr Leu Trp Glu Ile Gln Asn Lys Leu Lys	
335 340 345	
ctt taaagttgcc taaaactacc attccgaaat tattgaagag atcatagata	1266
Leu	
caggattata taacgaaatt ttgaataaac ttgaattcct aaaagatgga aacaggaaag	1326
taggtagagt gatttttccta tttatttagt cctccagctc ttttattgag catccacgtg	1386
ctggacgata cttattttaca attcckaagt atttttggtta cctctgatgt agcagcactt	1446
gccatgttat atatatgttag ttgrmatttg gttcccaaaa agtaggatgt aggtatttat	1506
tgtgttctag aaattccgac tcttttcatt agatatatgc tatttcctttc attcttgctg	1566
gtttatacct atgttcattt atattgctgta aaaaagtagt agcttccttct acaatgtaaa	1626
aataaatgta catacaaaaa aaaaaamcmc	1656

<210> 308  
<211> 517  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 48..164

<221> sig\_peptide  
<222> 48..89  
<223> Von Heijne matrix  
score 4  
seq YYMVCLFFRLIFS/EH

<221> polyA\_signal  
<222> 482..487

<221> polyA\_site  
<222> 505..517

<400> 308  
aggagatagc ctcgtagaaa tgacaaccac aatgttaata ctaacat atg tat tac 56  
Met Tyr Tyr  
atg gtt tgt ttg ttc ttt cgc tta ata ttt tca gag cac cta cct att 104  
Met Val Cys Leu Phe Phe Arg Leu Ile Phe Ser Glu His Leu Pro Ile  
-10 -5 1 5  
ata ggc act gtc act tct cac aaa act ggg aca cta act gtt tat cca 152  
Ile Gly Thr Val Thr Ser His Lys Thr Gly Thr Leu Thr Val Tyr Pro  
10 15 20  
aca tct gct ggc taaataaaga catgatcttc accttttggg attgttaatt 204  
Thr Ser Ala Gly  
25  
taaaatgggtt ccataagagc aatgcaaaga cagagatatt tggcagcact gcagctggtg 264  
attttatatgg ctcttcacaa ggtgttattt tggggatatca aggtatggat gcttaaatca 324  
gctgcaggaa gtaagaaaga agaaaaaagg agtgataaag ataaaaaaaa atcaaccttg 384  
gtccttccac caaaacccat taatttccat atcatcatct gcataararg gaaaattcct 444  
acwtgaccag gttactgcaa ggatktkaat tttgaatatt aaaatattat mcmcaattgg 504  
aaaaaaaaaa aaa 517

<210> 309  
<211> 405  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 185..334

<221> sig\_peptide  
<222> 185..295  
<223> Von Heijne matrix  
score 5.90000009536743

seq LSYASSALSPCLT/AP

<221> polyA\_signal

<222> 355..360

<221> polyA\_site

<222> 392..405

<400> 309

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atcaccttct tctccatcct tstctggggc agtccccarc ccagtccttc tcctgacctg      60
cccagcccaa gtcagccttc agcacgcgct tttctgcaca cagatattcc aggcctacct      120
ggcattccag gacctccgma atgatgctcc agtcccttac aagcgcttcc tggatgaggg      180
tggc atg gtg ctg acc acc ctc ccc ttg ccc tct gcc aac agc cct gtg      229
    Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val
          -35                -30                -25
aac atg ccc acc act ggc ccc aac agc ctg agt tat gct agc tct gcc      277
Asn Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala
          -20                -15                -10
ctg tcc ccc tgt ctg acc gct cca aag tcc ccc cga ctt gct atg atg      325
Leu Ser Pro Cys Leu Thr Ala Pro Lys Ser Pro Arg Leu Ala Met Met
          -5                1                5                10
cct gac aac taaatatacct tatccaaatc aataaarwra raatcctccc      374
Pro Asp Asn
cccaraaggg tttctaaaaa caaaaaaaaaa a      405

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<210> 310

<211> 1087

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 195..347

<221> sig\_peptide

<222> 195..272

<223> Von Heijne matrix

score 7.09999990463257

seq LASLQWSLTAWC/GS

<221> polyA\_signal

<222> 1037..1042

<221> polyA\_site

<222> 1071..1082

<400> 310

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aaagtgtaga acacggacct ctgagttatg ctcttgagag gtgccaaagc tgggctgttt      60
acctacctta tccacagagc tctgaaagtc aagccagaaa ggaaggattc caaattcttg      120
gaattttatc tagaaaagaa gactaagcag cttttgttct tctgtgaccc agttgctggc      180
ccaagacatg gaca atg acc ccc tgg tgt ttg gcg tgt ctg ggg agg agg      230
          Met Thr Pro Trp Cys Leu Ala Cys Leu Gly Arg Arg
          -25                -20                -15
cct ctc gct tct ttg cag tgg agc ctg aca ctg gcg tgg tgt ggc tcc      278
Pro Leu Ala Ser Leu Gln Trp Ser Leu Thr Leu Ala Trp Cys Gly Ser

```

```

-10          -5          1
ggc agc cac tgg aca gag aga cca akt cag akt tca ccg tgg akt tct    326
Gly Ser His Trp Thr Glu Arg Pro Xaa Gln Xaa Ser Pro Trp Xaa Ser
      5          10          15
ctg tca gcg acc acc agg ggg tgatcacacg gaaggtgaac atccaggtcg    377
Leu Ser Ala Thr Thr Arg Gly
      20          25
gggatgtgaa tgacaacgcg cccacatttc acaatcagcc ctacagcgtc cgcattccctg    437
araatacacc agtgggggacg cccatcttca togtgaatgc cacagacccc gacttgggggg    497
cagggggcag cgtcctctac tcttccagc cccctccca attcttcgcc attgacagcg    557
cccgcggtat cktcacagtg atccgggagc tggactacga taccacrcmg gcctaccagc    617
tcwcggtcwa cgccacagat caagacaara ccaggcctct gtccaccstg gccaaacttg    677
ccatcatcat cacagatgtc caggacatgg acccatctt catcaacctg ccttacagca    737
ccaacatcta cgagcattct cctccgggca cgacggtgcg catcatcacc gccatagacc    797
aggataaagg acgtccccgg ggcattggct acaccatcgt ttcagggcat ctgtgtttac    857
aagaacccaa gatctctcag gagctcagga aaaggggctt gctgtgaggc tcagggttcc    917
catggacatt ctgagctgac cctcctcagc attggatctc ctggctcagg aactaggaac    977
gaagcttgga tgttttctcc tttcctacag catctgtatt catttcctat agttgccata   1037
ataaaatgcc actaacttag tggcttaaaa accaaaaaaa aaaaaccctt   1087
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<210> 311
<211> 916
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 90..815

<221> sig_peptide
<222> 90..179
<223> Von Heijne matrix
      score 13.1999998092651
      seq LLLLSTLVIPSAA/AP

<221> polyA_signal
<222> 883..888

<221> polyA_site
<222> 905..916
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```

<400> 311
aaaacagtac gtgggcgggc ggaatccggg agtccggtga cccgggctgt ggtctagcat    60
aaaggcggag ccagaagaag gggcgggggt atg gga gaa gcc tcc cca cct gcc    113
                        Met Gly Glu Ala Ser Pro Pro Ala
                        -30          -25
ccc gca agg cgg cat ctg ctg gtc ctg ctg ctg ctc ctc tct acc ctg    161
Pro Ala Arg Arg His Leu Leu Val Leu Leu Leu Leu Leu Ser Thr Leu
      -20          -15          -10
gtg atc ccc tcc gct gca gct cct atc cat gat gct gac gcc caa gag    209
Val Ile Pro Ser Ala Ala Pro Ile His Asp Ala Asp Ala Gln Glu
      -5          1          5          10
agc tcc ttg ggt ctc aca ggc ctc cag agc cta ctc caa ggc ttc agc    257
Ser Ser Leu Gly Leu Thr Gly Leu Gln Ser Leu Leu Gln Gly Phe Ser
      15          20          25
```



cga ctt ttc ctg aaa ggt aac ctg ctt cgg ggc ata gac agc tta ttc	305
Arg Leu Phe Leu Lys Gly Asn Leu Leu Arg Gly Ile Asp Ser Leu Phe	
30 35 40	
tct gcc ccc atg gac ttc cgg ggc ctc cct ggg aac tac cac aaa gag	353
Ser Ala Pro Met Asp Phe Arg Gly Leu Pro Gly Asn Tyr His Lys Glu	
45 50 55	
gag aac cag gag cac cag ctg ggg aac aac acc ctc tcc agc cac ctc	401
Glu Asn Gln Glu His Gln Leu Gly Asn Asn Thr Leu Ser Ser His Leu	
60 65 70	
cag atc gac aag atg acc gac aac aag aca gga gag gtg ctg atc tcc	449
Gln Ile Asp Lys Met Thr Asp Asn Lys Thr Gly Glu Val Leu Ile Ser	
75 80 85 90	
gag aat gtg gtg gca tcc att caa cca vcg gag ggg anc ttc gag ggt	497
Glu Asn Val Val Ala Ser Ile Gln Pro Xaa Glu Gly Xaa Phe Glu Gly	
95 100 105	
gat ttg aag gth ccc agg atg gag gar aag gag gcc ctg gta ccc mtc	545
Asp Leu Lys Val Pro Arg Met Glu Glu Lys Glu Ala Leu Val Pro Xaa	
110 115 120	
car aag gcc acg gac agc ttc cac aca gaa ctc cat ccc cgg gtg gcc	593
Gln Lys Ala Thr Asp Ser Phe His Thr Glu Leu His Pro Arg Val Ala	
125 130 135	
ttc tgg atc att aag ctg cca cgg cgg agg tcc cac cag gat gcc ctg	641
Phe Trp Ile Ile Lys Leu Pro Arg Arg Arg Ser His Gln Asp Ala Leu	
140 145 150	
gag ggc ggc cac tgg ctc anc gar aag cga cac cgc ctg cag gcc atc	689
Glu Gly Gly His Trp Leu Xaa Glu Lys Arg His Arg Leu Gln Ala Ile	
155 160 165 170	
cgg gat gga ctc cgc aag ggg acc cac aag gac rtc cta daa rag ggg	737
Arg Asp Gly Leu Arg Lys Gly Thr His Lys Asp Xaa Leu Xaa Xaa Gly	
175 180 185	
acc gar agc tcc tcc cac tcc agg ctg tcc ccc cga aar amm cac tta	785
Thr Glu Ser Ser Ser His Ser Arg Leu Ser Pro Arg Lys Xaa His Leu	
190 195 200	
ctg tac atc ctc arg ccc tct cgg cag ctg targgggtggg gaccgggggar	835
Leu Tyr Ile Leu Xaa Pro Ser Arg Gln Leu	
205 210	
macctgctg tagcccccat caraccctgc cccaagcacc atatggaaat aaagttcttt	895
cttacatcca aaaaaaaaaa a	916

<210> 312

<211> 583

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 52..513

<221> sig\_peptide

<222> 52..231

<223> Von Heijne matrix

score 4

seq LVRRTLLVAALRA/WM

<221> polyA\_signal

<222> 553..558

<221> polyA\_site

<222> 572..583

<400> 312

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aaggaaacag caaccagagg gagatgatca cctgaaccac tgctccaaac c atg ggc      57
                                     Met Gly
                                     -60
agt aaa tgc tgt aaa ggt ggt cca gat gaa gat gca gta gaa aga cag      105
Ser Lys Cys Cys Lys Gly Gly Pro Asp Glu Asp Ala Val Glu Arg Gln
      -55                    -50                    -45
agg cgg cag aag ttg ctt ctt gca caa ctg cat cac aga aaa agg gtg      153
Arg Arg Gln Lys Leu Leu Leu Ala Gln Leu His His Arg Lys Arg Val
      -40                    -35                    -30
aar gca gct ggg cag atc cag gcc tgg tgg cgt ggg gtc ctg gtg cgc      201
Lys Ala Ala Gly Gln Ile Gln Ala Trp Trp Arg Gly Val Leu Val Arg
      -25                    -20                    -15
agg acc ctg ctg gtt gct gcc ctc agg gcc tgg atg att cag tgc tgg      249
Arg Thr Leu Leu Val Ala Ala Leu Arg Ala Trp Met Ile Gln Cys Trp
      -10                    -5                    1                    5
tgg agg acg ttg gtg cag aga cgg atc cgt cag cgg cgg cag gcc ctg      297
Trp Arg Thr Leu Val Gln Arg Arg Ile Arg Gln Arg Arg Gln Ala Leu
      10                    15                    20
ctt ggg gtc tac gtc atc cag gag cag gcg gcg gtc aag ctc cag tcc      345
Leu Gly Val Tyr Val Ile Gln Glu Gln Ala Ala Val Lys Leu Gln Ser
      25                    30                    35
tgc atc cgc atg tgg cag tgc cgg caa tgt tac cgc caa atg tgc aat      393
Cys Ile Arg Met Trp Gln Cys Arg Gln Cys Tyr Arg Gln Met Cys Asn
      40                    45                    50
gct ctc tgc ttg ttc cag gtc cca aaa agc agc ctt gcc ttc caa act      441
Ala Leu Cys Leu Phe Gln Val Pro Lys Ser Ser Leu Ala Phe Gln Thr
      55                    60                    65                    70
gat ggc ttt tta cag gtc caa tat gca atc cct tca aag cag cca gag      489
Asp Gly Phe Leu Gln Val Gln Tyr Ala Ile Pro Ser Lys Gln Pro Glu
      75                    80                    85
ttc cac att gaa atc cta tca atc tgaaaggcct ggggcatgga gaacaggctg      543
Phe His Ile Glu Ile Leu Ser Ile
      90
cactacccta ataatgtct gaccaggtaa aaaaaaaaaa      583
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<210> 313

<211> 697

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 172..438

<221> sig\_peptide

<222> 172..354

<223> Von Heijne matrix

score 4.69999980926514

seq LLPCNLHCSWLHS/SP

<221> polyA\_signal  
<222> 682..687

<221> polyA\_site  
<222> 685..697

<400> 313

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agattggctg ggcagatggg ctgactggct gggcagatgg gtgggtgagt tccctctccc      60
cagagccatc ggccaggtac caaagctcag ctgtatggat tcccaacagg aggacctgcg      120
cttccctggg acccattggt gtactggatt aacaagcgac ggcgctacgg c atg aat      177
                                     Met Asn
                                     -60
gca gcc atc aac acg ggc cct gcc cct gct gtc acc aag act gag act      225
Ala Ala Ile Asn Thr Gly Pro Ala Pro Ala Val Thr Lys Thr Glu Thr
                                     -55                                     -50                                     -45
gag gtc cag aat cca gat gtt ctg tgg gat ttg gac atc ccc gaa gcc      273
Glu Val Gln Asn Pro Asp Val Leu Trp Asp Leu Asp Ile Pro Glu Ala
                                     -40                                     -35                                     -30
agg agc cat gct gac caa gac agc aac ccc aag gcg gaa gcc ctg ctc      321
Arg Ser His Ala Asp Gln Asp Ser Asn Pro Lys Ala Glu Ala Leu Leu
                                     -25                                     -20                                     -15
ccc tgc aac ctg cac tgc agc tgg ctc cac agc agc ccc agg cca gat      369
Pro Cys Asn Leu His Cys Ser Trp Leu His Ser Ser Pro Arg Pro Asp
                                     -10                                     -5                                     1                                     5
ccc cat tcc cac ttc cca tct ktc agg agg tgc cct ttg ccc cac cct      417
Pro His Ser His Phe Pro Ser Xaa Arg Arg Cys Pro Leu Pro His Pro
                                     10                                     15                                     20
tgt gca acc tac ccc ccs kgc tgaaccactc tgtctcctat cctttggcca      468
Cys Ala Thr Tyr Pro Pro Xaa
                                     25
cctgtcctga aaggaatggt ctcttccatt ccctcctgaa tctggcccag gaagaccata      528
ccttcaatgy caagcctttt ccttcaaaac tgtagcctcc tctcactgaa ggtgggagct      588
gcaggaatca ggtgcagagt aggaaatgga actaacctca ggaaggtggt attgacagag      648
gtcaggaccc acctggatgt catgctatga aacattaaaa gaaaaaaaaa      697

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<210> 314  
<211> 803  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 148..366

<221> sig\_peptide  
<222> 148..225  
<223> Von Heijne matrix  
score 5.5  
seq LFTLLFLIMLVLK/LD

<221> polyA\_signal  
<222> 770..775

<221> polyA\_site

<222> 792..803

<400> 314

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aaatggggggg aaaagggcgg aaaaggacaa ggatccaaac tggcgaattt gctgatcttc      60
gcgtccctct cgcgtttccg gccggcagcg ctgccagggt atatttcctt ttttccgac      120
ctgcaacagc ctcttttaaac tgttttaa atg aga atg tcc ttg gct cag aga gta      174
                               Met Arg Met Ser Leu Ala Gln Arg Val
                               -25                -20

cta ctc acc tgg ctt ttc aca cta ctc ttc ttg atc atg ttg gtg ttg      222
Leu Leu Thr Trp Leu Phe Thr Leu Leu Phe Leu Ile Met Leu Val Leu
      -15                -10                -5

aaa ctg gat gag aaa gca cct tgg aac tgg ttc ctc ata ttc att cca      270
Lys Leu Asp Glu Lys Ala Pro Trp Asn Trp Phe Leu Ile Phe Ile Pro
      1                5                10                15

gtc tgg ata ttt gat act atc ctt ctt gtc ctg ctg att gtg aaa atg      318
Val Trp Ile Phe Asp Thr Ile Leu Leu Val Leu Leu Ile Val Lys Met
      20                25                30

gct ggg cgg tgt aag tct ggc ttt gac ctc gac atg gat cac aca ata      366
Ala Gly Arg Cys Lys Ser Gly Phe Asp Leu Asp Met Asp His Thr Ile
      35                40                45

taaaaaaaaa aacctggtac ctcatcgac tgktacttaa attasccttc tgcctcgac      426
tctgtgctaa actggaacag ttactacca tgaatctatc ctatgtcttc attcctttat      486
gggccttgct ggctggggct ttaacagAAC tcggatataa tgtctttttt gtgaaagact      546
gacttctaag tacatcatct cctttctatt gctgttcaac aagttaccat taaagtgttc      606
tgaatctgtc aagcttcaag aataccagag aactgagggg aaataccaaa tgtagtttta      666
tactacttcc ataaaacagg attggtgaat cacggacttc tagtcaacct acagcttaat      726
tattcagcat ttgagttatt gaaatcctta ttatctctat gtaaataaag tttgttttgg      786
aacctcaaaaa aaaaaaa      803
```

<210> 315

<211> 823

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 175..336

<221> sig\_peptide

<222> 175..276

<223> Von Heijne matrix

score 3.70000004768372

seq SVLNVGHLLFSSA/CS

<221> polyA\_site

<222> 812..823

<400> 315

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aaggcgcgcg cgaccggcgg ctctttggcg cggattaggg ggtctcgcg agggagtcac      60
caagctttgg tgtatgtgtt ggccggttct gaagtcttga agaagctctg ctgaggaaga      120
ccaaagcagc actcgttgcc aattagggaa tggaccgttt gggttccttt agca atg      177
                               Met
atc cct ctg ata agc cac ctt gcc gag gct gct cct cct acc tca tgg      225
Ile Pro Leu Ile Ser His Leu Ala Glu Ala Ala Pro Pro Thr Ser Trp
      -30                -25                -20
```

```

agc ctt ata tca agt gtg ctg aat gtg ggc cac ctc ctt ttt tcc tct      273
Ser Leu Ile Ser Ser Val Leu Asn Val Gly His Leu Leu Phe Ser Ser
      -15                      -10                      -5
gct tgc agt gtt tca ctc gag gct ttg agt aca aga aac atc aaa gcg      321
Ala Cys Ser Val Ser Leu Glu Ala Leu Ser Thr Arg Asn Ile Lys Ala
      1                      5                      10                      15
atc ata ctt atg aaa taatggcttc agattttcct gtccttgatc ccagctggac      376
Ile Ile Leu Met Lys
      20
tgctcaagaa raaatggccc ttttagaasc tgtgatggac tgtggctttg gaaattggca      436
ggatgtagcc aatcaaatgt gcaccaarac caaggaggag tgtgagaagc actatatgaa      496
gcatttcac aataaccctc tgtttgcac trscctgctg aacctgaaac aascagrgga      556
agcaaaaact gctgacacag ccattccatt tcaactctaca ratgaccctc cccgacckac      616
ctttgactcc ttgctttctc gggacatggc cgggtacwtg ccmgctcgag cagatttcac      676
tgaggaatth gacaattatg cagaatggga cttgagagac attgattttg ttgaagatga      736
ctcgacatt ttacatgctc tgaagatggc tgtggtagat atctatcatt ccagggttaa      796
ggagagacaa agacgaaaaa aaaaaaa
      823

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<210> 316
<211> 823
<212> DNA
<213> Homo sapiens
<220>
<221> CDS
<222> 191..553
<221> sig_peptide
<222> 191..304
<223> Von Heijne matrix
      score 5.69999980926514
      seq LAFLSCLAFLVLD/TQ
<221> polyA_signal
<222> 766..771
<221> polyA_site
<222> 804..817

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<400> 316
aactctgcag ggcctccaag gccaggcttc agggctggga ctcagtcctg aggcactggg      60
gagccatgag gggctgtggc agggaggggc aggggtgtga aagactcccc tggggccatg      120
gtggagatgt gctgaggtct tctccctgat cgtcttctcc tccctgctga ccgacggcta      180
ccagaackag atg gag tct ccg cag ctc cac tgc att ctc aac agc aac      229
      Met Glu Ser Pro Gln Leu His Cys Ile Leu Asn Ser Asn
      -35                      -30
agc gtg gcc tgc agc ttt gcc gtg gga gcc ggc ttc ctg gcc ttc ctc      277
Ser Val Ala Cys Ser Phe Ala Val Gly Ala Gly Phe Leu Ala Phe Leu
      -25                      -20                      -15                      -10
agc tgc ctg gcc ttc ctc gtc ctg gac aca cag gag acc cgc att gcc      325
Ser Cys Leu Ala Phe Leu Val Leu Asp Thr Gln Glu Thr Arg Ile Ala
      -5                      1                      5
ggc acc cgc ttc aag aca gcc ttc cag ctc ctg gac ttc atc ctg gct      373
Gly Thr Arg Phe Lys Thr Ala Phe Gln Leu Leu Asp Phe Ile Leu Ala
      10                      15                      20

```

```

gtt ctc tgg gca gtt gtc tgg ttc atg ggt ttc tgc ttc ctg gcc aac      421
Val Leu Trp Ala Val Val Trp Phe Met Gly Phe Cys Phe Leu Ala Asn
    25                      30                      35
caa tgg cag cat tcg ccg ccc aaa gar kkc ctc ctg ggg agc agc agt      469
Gln Trp Gln His Ser Pro Pro Lys Glu Xaa Leu Leu Gly Ser Ser Ser
    40                      45                      50                      55
gcc cag gca gcc atc ggc stt cac ctt ctt ctc cat cct tgt ctg gat      517
Ala Gln Ala Ala Ile Gly Xaa His Leu Leu Leu His Pro Cys Leu Asp
    60                      65                      70
att cca rgc cta cct ggc akk cca gga cct ccg aaa tgatgctcca      563
Ile Pro Xaa Leu Pro Gly Xaa Pro Gly Pro Pro Lys
    75                      80
gtcccttacm arcgcttcct ggatgaaggt ggcattggtgs kkaacaccct ccccttgccc      623
tctgccaaca gcctgtgaac atgcccacca ctggcccca cagcctgagt tatgctagct      683
ctgccctgtc cccctgtctg accgctcmaa agtccccccg gcttgctatg atgcctgaca      743
actaaatata cttatccaaa tcaataaaga gagaatcctc cctccagaag ggtttctaaa      803
aacaacaaaaa aaaahncctt      823

```

<210> 317

<211> 1112

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 106..603

<221> sig\_peptide

<222> 106..216

<223> Von Heijne matrix

score 4.30000019073486

seq LWEKLTLLSPGIA/VT

<221> polyA\_site

<222> 1102..1112

<400> 317

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agcgattgcg aatcctccgc tgaggtgatt tggatatccc tagaacgttg agggcacgag      60
tcgggtcctg agaccagggtc ctcagccagc agagccacgt tcctt atg agc acc gtg      117
                                Met Ser Thr Val
                                -35
ggt tta ttt cat ttt cct aca cca ctg acc cga ata tgc ccg gcg cca      165
Gly Leu Phe His Phe Pro Thr Pro Leu Thr Arg Ile Cys Pro Ala Pro
    -30                      -25                      -20
tgg gga ctc cgg ctt tgg gag aag ctg acg ttg tta tcc cca gga ata      213
Trp Gly Leu Arg Leu Trp Glu Lys Leu Thr Leu Leu Ser Pro Gly Ile
    -15                      -10                      -5
gct gtc act ccg gtc cag atg gca ggc aag aag gac tac cct gca ctg      261
Ala Val Thr Pro Val Gln Met Ala Gly Lys Lys Asp Tyr Pro Ala Leu
    1                      5                      10                      15
ctt tcc ttg gat gag aat gaa ctc gaa gag cag ttt gtg aaa gga cac      309
Leu Ser Leu Asp Glu Asn Glu Leu Glu Glu Phe Val Lys Gly His
    20                      25                      30
ggt cca ggg ggc cag gca acc aac aaa acc agc aac tgc gtg gtg ctg      357
Gly Pro Gly Gly Gln Ala Thr Asn Lys Thr Ser Asn Cys Val Val Leu

```

```

      35              40              45
aar mac atc ccc tca ggc atc gtt gta aag tgc cat cag aca aga tca      405
Lys Xaa Ile Pro Ser Gly Ile Val Val Lys Cys His Gln Thr Arg Ser
      50              55              60
gtt gat cag aac aga aag cta gct cgg aaa atc cta caa gag aaa gta      453
Val Asp Gln Asn Arg Lys Leu Ala Arg Lys Ile Leu Gln Glu Lys Val
      65              70              75
rat gtt ttc tac aat ggt gaa aac agt cct gtt cac aaa gaa aaa cga      501
Xaa Val Phe Tyr Asn Gly Glu Asn Ser Pro Val His Lys Glu Lys Arg
      80              85              90              95
gaa gcg gcg aag aaa aaa car gaa agg aaa aaa aga gca aag gaa acc      549
Glu Ala Ala Lys Lys Lys Gln Glu Arg Lys Lys Arg Ala Lys Glu Thr
      100              105              110
ctg gaa aaa aag aas ctm ctt aaa raa ctg tgg gag tca agt aaa aag      597
Leu Glu Lys Lys Xaa Leu Leu Lys Xaa Leu Trp Glu Ser Ser Lys Lys
      115              120              125
gtc cac tgagaaaaga attagagatt ccaactgaca gaatctgcca gaagctccca      653
Val His
gggaataatg gtggcgagtt ccatcaccag cattattata gtgcttcaaa agaaatattt      713
ttgatgaact taaaagacaa caaatttatt taaatggtgc actaaactgt agtgaacaga      773
gacatgcacg attcaagaat aaaactoggc cgggcacggt ggacggtgcc tcacatctgt      833
aatcccagca ctttgggagg ccgaggcggg cggatcactt gaggtcagga gtttgagacc      893
agcctggcca acatggtgaa accccgtctc tactaaaaat acaaaaaatt agccaggcat      953
gggtggcgggc acctgtaatc ccagctactc gggaggccga ggcaggagaa ttgctgaac      1013
ctgggaggcg gaggttgag tgagctgaga tcgcgccact gcactcaagc ctgggcaaca      1073
cctgggtgac agagcaagac cccatcycaa aaaaaaaaaa      1112

<210> 318
<211> 1623
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 47..586

<221> sig_peptide
<222> 47..124
<223> Von Heijne matrix
      score 6.30000019073486
      seq GVGLVTLLGLAVG/SY

<221> polyA_signal
<222> 1583..1588

<221> polyA_site
<222> 1614..1623

<400> 318
agggatctgt cggcttgtca ggtggtggag gaaaaggcgc tccgtc atg ggg atc      55
                                         Met Gly Ile
                                         -25
cag acg agc ccc gtc ctg ctg gcc tcc ctg ggg gtg ggg ctg gtc act      103
Gln Thr Ser Pro Val Leu Leu Ala Ser Leu Gly Val Gly Leu Val Thr
      -20              -15              -10

```

ctg ctc ggc ctg gct gtg ggc tcc tac ttg gtt cgg agg tcc cgc cgg	151
Leu Leu Gly Leu Ala Val Gly Ser Tyr Leu Val Arg Arg Ser Arg Arg	
-5 1 5	
cct cag gtc act ctc ctg gac ccc aat gaa aag tac ctg cta cga ctg	199
Pro Gln Val Thr Leu Leu Asp Pro Asn Glu Lys Tyr Leu Leu Arg Leu	
10 15 20 25	
cta gac aag acg act gtg agc cac aac acc aag agg ttc cgc ttt gcc	247
Leu Asp Lys Thr Thr Val Ser His Asn Thr Lys Arg Phe Arg Phe Ala	
30 35 40	
ctg ccc acc gcc cac cac act ctg ggg ctg cct gtg ggc aaa cat atc	295
Leu Pro Thr Ala His His Thr Leu Gly Leu Pro Val Gly Lys His Ile	
45 50 55	
tac ctc tcc acm mga att gat ggc agc ctg gtc atc agg cca tac act	343
Tyr Leu Ser Thr Arg Ile Asp Gly Ser Leu Val Ile Arg Pro Tyr Thr	
60 65 70	
cct gtc acc agt gat gag gat caa ggc tat gtg gat ctt gtc mtc aag	391
Pro Val Thr Ser Asp Glu Asp Gln Gly Tyr Val Asp Leu Val Xaa Lys	
75 80 85	
gtc tac ctg aag ggt gtg cac ccc aaa ttt cct gag gga ggg aar atg	439
Val Tyr Leu Lys Gly Val His Pro Lys Phe Pro Glu Gly Gly Lys Met	
90 95 100 105	
tct cak tac ctg gat asc ctg aaa gtt ggg gat btg gtg gaa ttt csg	487
Ser Xaa Tyr Leu Asp Xaa Leu Lys Val Gly Asp Xaa Val Glu Phe Xaa	
110 115 120	
ggg cca agc ggg ttg ctc act tac act gga aaa ggg cat ttt aac att	535
Gly Pro Ser Gly Leu Leu Thr Tyr Thr Gly Lys Gly His Phe Asn Ile	
125 130 135	
cag ccc aac aag aat ctc cac cag aac ccc gag tgg cga aga aac tgg	583
Gln Pro Asn Lys Asn Leu His Gln Asn Pro Glu Trp Arg Arg Asn Trp	
140 145 150	
gaa tgattgccgg cgggacagga atcaccccaa tgctacagct gatccgggcc	636
Glu	
atcctgaaag tccctgaaga tccaaccag tgctttctgc tttttgccaa ccagacagaa	696
aaggatatca tcttgcgga ggacttagag gaactgcagg cccgctatcc caatcgcttt	756
aagctctggt tcaactctga tcatcccca aaagrttggg cctacagcaa gggctttgtg	816
actgccgacw tgatccggga acacctgccc gctccagggg atgatgtgct ggtactgctt	876
tggtgggcmc cccaatggt gcagctggcc tgccatcca acttggaaca actgggctac	936
tcacaaaaga tgcgattcac ctactgagca tcctccagct tccctggtgc tgttcgctgc	996
agttgttccc catcagtact caagcactak aagccttagr ktcctktcct cagagtttca	1056
ggtttttttca gttrsatcka gagctgaaat ctggatagta cctgcaggaa caatattcct	1116
gtagccatgg aagagggcca aggctcagtc actccttgga tggcctccta aatctccccg	1176
tggcaacagg tccaggagag gcccatggag cagtctcttc catggagtaa gaaggaagg	1236
agcatgtacg cttggtccaa gattggctag ttccttgata gcatcttact ctacacttct	1296
ttgtgtctgt gatgaaagga acagtctgtg caatgggttt tacttaaact tcactgttca	1356
acctatgagc aaatctgtat gtgtgagtat aagttgagca tagcatactt ccagaggtgg	1416
tcttatggag atggcaagaa aggaggaaat gatttcttca gatctcaaag gagtctgaaa	1476
tatcatattt ctgtgtgtgt cdctctcagc ccctgcccad gctagaggga wacagctact	1536
gataatcgaa aactgctgtt tgtgggcarg aacccttggc tgtgcaaata atggggctga	1596
ngccctgtgt gatattgaaa aaaaaa	1623

<210> 319

<211> 526

<212> DNA

<213> Homo sapiens



<220>

<221> CDS

<222> 99..371

<221> sig\_peptide

<222> 99..290

<223> Von Heijne matrix

score 3.79999995231628

seq LFIVVCVICVTLN/FP

<221> polyA\_signal

<222> 491..496

<221> polyA\_site

<222> 513..524

<400> 319

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attggattag tagaattgct tttgtcattc cattgttttc atatatttgt ttgggacatt      60
ttactttttt ctgttaacgc ttaccctagr aattagaa atg aca cca cgt att ctt      116
                               Met Thr Pro Arg Ile Leu
                               -60
agc gaa gtc cag ttt tca gca ttt tgt cct tat tgg aca ata gca agg      164
Ser Glu Val Gln Phe Ser Ala Phe Cys Pro Tyr Trp Thr Ile Ala Arg
                               -55                               -50                               -45
ata tta gaa cgt gtt ggt tcc gcg tgc ttc cgt ctt gag tta tgt gct      212
Ile Leu Glu Arg Val Gly Ser Ala Cys Phe Arg Leu Glu Leu Cys Ala
                               -40                               -35                               -30
gct att gtc gga tat ttt gtc tta gat gta cgt act ttc ctg ttc att      260
Ala Ile Val Gly Tyr Phe Val Leu Asp Val Arg Thr Phe Leu Phe Ile
                               -25                               -20                               -15
gtg gta tgt gta att tgc gtt act ttg aat ttt cca cgt ttt tac ttt      308
Val Val Cys Val Ile Cys Val Thr Leu Asn Phe Pro Arg Phe Tyr Phe
10                               -5                               1                               5
ctt tgt ctc tca tca ctt acc gct ttt ggg acc ccc ccc atc ggg gtt      356
Leu Cys Leu Ser Ser Leu Thr Ala Phe Gly Thr Pro Pro Ile Gly Val
                               10                               15                               20
cac att ccc tct ccc tararcacac tcccttgat ttcctcradt ggggtctgct      411
His Ile Pro Ser Pro
25
gcggtgaagc tttcccatTT tatgtgcaga ttattttcag agggtatata gaattcaggc      471
agctgtttcg ttgtagcaca ttaaaaatat tttcccactt caaaaaaaaaa aaacc      526

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<210> 320

<211> 989

<212> DNA

<213> Homo sapiens

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<222> 44..814

<221> sig\_peptide

<222> 44..112

<223> Von Heijne matrix

score 8.30000019073486

seq VRLLLXLLLLLLIA/LE

<221> polyA site

<222> 978. . 989

<400> 320

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								-20	
tcc	ctg	act	tct	agc	cct	gtg	cgc	ctt	103
Ser	Leu	Thr	Ser	Ser	Pro	Val	Arg	Leu	
				-15				-10	
cta	ata	gcc	ttg	gag	atc	atg	gtt	ggg	151
Leu	Ile	Ala	Leu	Glu	Ile	Met	Val	Gly	
			1			5			
ttc	act	ata	aaa	tca	ttg	tcc	aga	cct	199
Phe	Thr	Ile	Lys	Ser	Leu	Ser	Arg	Pro	
	15				20				
cat	gtc	ttc	ttg	aat	aaa	aat	ctt	ttc	247
His	Val	Phe	Leu	Asn	Lys	Asn	Leu	Phe	
				35				40	
aac	atg	gtc	aaa	cct	ctg	ggc	ctc	ctg	295
Asn	Met	Val	Lys	Pro	Leu	Gly	Leu	Leu	
				50				55	
agc	act	tgg	gga	gaa	ttg	acc	caa	acg	343
Ser	Thr	Trp	Gly	Glu	Leu	Thr	Gln	Thr	
			65				70		
ctc	agg	atg	ctc	ctt	tgt	gac	atc	aaa	391
Leu	Arg	Met	Leu	Leu	Cys	Asp	Ile	Lys	
			80			85			
cct	tcc	act	ctg	caa	gtc	kar	atk	ttt	439
Pro	Ser	Thr	Leu	Gln	Val	Xaa	Xaa	Phe	
					100				
tgc	act	ggg	gca	tcc	tgg	cag	ttc	gcc	487
Cys	Thr	Gly	Ala	Ser	Trp	Gln	Phe	Ala	
				115				120	
ctc	ttt	gac	gca	atg	aac	atg	acc	tgg	535
Leu	Phe	Asp	Ala	Met	Asn	Met	Thr	Trp	
				130				135	
agt	wag	atc	aag	gag	aca	tgg	aag	aaa	583
Ser	Xaa	Ile	Lys	Glu	Thr	Trp	Lys	Lys	
			145					150	
ttc	agg	aag	ctc	tca	aar	gga	gac	tgc	631
Phe	Arg	Lys	Leu	Ser	Lys	Gly	Asp	Cys	
			160			165			
tta	ggg	cac	tgg	gaa	gca	atg	cca	raa	679
Leu	Gly	His	Trp	Glu	Ala	Met	Pro	Xaa	
					180			185	
gct	tca	raw	atc	cac	tgg	tct	tct	tct	727
Ala	Ser	Xaa	Ile	His	Trp	Ser	Ser	Ser	
				195				200	
atc	ctg	ggg	gca	ttc	atc	ctg	tta	vtt	775
Ile	Leu	Gly	Ala	Phe	Ile	Leu	Leu	Xaa	
				210				215	
tgt	gtc	tgg	tgg	caa	aat	ggc	ara	ara	824
Cys	Val	Trp	Trp	Gln	Asn	Gly	Xaa	Xaa	
				225				230	

gcggcgcaaa attgttcacc tgtggctctc gatcgctgac agccttggct cccactgctg 884  
 tgtgttcctt gagtcaagtg gaggcggagc ctgcaatgag cggaratcgc gcctctgcat 944  
 tccagtcttg gcaacagarc aagactccgt ctcaaaaaaa aaaaa 989

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 <223> Von Heijne matrix  
 score 6.69999980926514  
 seq LWPFLTWINPALS/IC

<221> polyA\_site  
 <222> 1006..1016

<400> 321  
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 Met Cys Pro Ser Leu Glu Glu Ala Pro Ser Val Lys Gly Thr Leu  
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 ccc tgc tca gga caa cag cag cct ttc ccg ttt gga gcc tca aac atc 95  
 Pro Cys Ser Gly Gln Gln Gln Pro Phe Pro Phe Gly Ala Ser Asn Ile  
 -45 -40 -35 -30  
 cca cta ctc ctg ggc agg agc aga aag gtg gct cga ggt gca ccg gtc 143  
 Pro Leu Leu Leu Gly Arg Ser Arg Lys Val Ala Arg Gly Ala Pro Val  
 -25 -20 -15  
 ctg tgg cca ttt ctc act tgg ata aac cct gca ctg tcc atc tgt gac 191  
 Leu Trp Pro Phe Leu Thr Trp Ile Asn Pro Ala Leu Ser Ile Cys Asp  
 -10 -5 1  
 ccc tta gga tcc tgc gga tgg cyw tgc cac acg gcc car gtc cct gcg 239  
 Pro Leu Gly Ser Cys Gly Trp Xaa Cys His Thr Ala Gln Val Pro Ala  
 5 10 15  
 ccc ctg car ttg cct act gcc tgt cct ccc ctc cca cat ggc acc cgg 287  
 Pro Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg  
 20 25 30 35  
 gct gta ggc ccc acg cca ggc ctc ctc cct gag gct gca gcc cca sgc 335  
 Ala Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Ala Pro Xaa  
 40 45 50  
 acg tgk ggg gca ctg tcc tca cgc agc agg cac tgg tca tgt tcc att 383  
 Thr Xaa Gly Ala Leu Ser Ser Arg Ser Arg His Trp Ser Cys Ser Ile  
 55 60 65  
 gtc arc tgc ctc cac ctg cac ara ctc ctg tct gtg gag acc aga arc 431  
 Val Xaa Cys Leu His Leu His Xaa Leu Leu Ser Val Glu Thr Arg Xaa  
 70 75 80  
 ttc cas aaa cat ctg ttg gtg ctg ctg gtg gct gtg gcc cat agt gtt 479  
 Phe Xaa Lys His Leu Leu Val Leu Leu Val Ala Val Ala His Ser Val  
 85 90 95  
 ctg gaa cca cct gcc ctg gtc cca aat gtg cag tgt gag atg tgc aca 527  
 Leu Glu Pro Pro Ala Leu Val Pro Asn Val Gln Cys Glu Met Cys Thr

100	105	110	115	
cac tca ggg ccc cgt gac ctg gaa gcc gca gtc gtg tcc cca gca cct				575
His Ser Gly Pro Arg Asp Leu Glu Ala Ala Val Val Ser Pro Ala Pro				
	120	125	130	
tgg gaa tgagcctgtc ctctgtgtga aggaggggggt gggttctcaaa ccactgactc				631
Trp Glu				
ttggtgctca ggagggggcct gctgctgtcc tgggcatggg gtgggtcattg ttcaagactg				691
aggcagactc agtctttgaa aggggtgcaga ggccaggcgc ggtgggtcac gcctgtaatt				751
ccagcacttt gggaggccaa ggtggacaga tcatgaggtc aggagttcga gaccagcctg				811
gccaatacgg tgaaaccgca tctctactaa rraatawcaw aaattagtcg ggcatgggtg				871
atgtgtgctt gtagtcccag ctactcatga ggyctgaggc agaagaatca cctgaatctg				931
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 score 3.79999995231628  
 seq RFLSLSAADGSDG/SH  
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 <222> 499..504  
 <221> polyA\_site  
 <222> 516..529

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	Met Val Leu
acg ctc gga gaa agt tgg ccg gta ttg gtg ggg agg agg ttt ctc agt	163
Thr Leu Gly Glu Ser Trp Pro Val Leu Val Gly Arg Arg Phe Leu Ser	
-25 -20 -15 -10	
ctg tcc gca gcc gac ggc agc gat ggc agc cac gac agc tgg gac gtg	211
Leu Ser Ala Ala Asp Gly Ser Asp Gly Ser His Asp Ser Trp Asp Val	
-5 1 5	
gag cgc gtc gcc gag tgg ccc tgg ctc tcc ggg acc att cga gct gtt	259
Glu Arg Val Ala Glu Trp Pro Trp Leu Ser Gly Thr Ile Arg Ala Val	
10 15 20	
tcc cac acc gac gtt acc aag aag gat ctg aag gtg tgt gtg gaa ttt	307
Ser His Thr Asp Val Thr Lys Lys Asp Leu Lys Val Cys Val Glu Phe	
25 30 35	
gak ggg gaa tct tgg agg aaa aga aga tgg ata gaa gtc tac agc ctt	355
Xaa Gly Glu Ser Trp Arg Lys Arg Arg Trp Ile Glu Val Tyr Ser Leu	
40 45 50 55	
cta agg aaa gca ttt tta gta aaa cat aat ttg gtt tta gct gaa cga	403

Leu Arg Lys Ala Phe Leu Val Lys His Asn Leu Val Leu Ala Glu Arg  
60 65 70  
aag tca cct gaa att tct tgg ggt taaccatctt tagttaaatg gaattttaat 457  
Lys Ser Pro Glu Ile Ser Trp Gly  
75  
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aaaaaaaaaa aa 529

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<221> sig\_peptide  
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score 5.69999980926514  
seq MLVLRSALETRALA/SR

<221> polyA\_signal  
<222> 1008..1013  
<221> polyA\_site  
<222> 1032..1042

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Met Leu Val Leu  
-10  
aga agc gcc ctg act cgg gcg ctg gcc tca cgg acg ctg gcg cct cag 104  
Arg Ser Ala Leu Thr Arg Ala Leu Ala Ser Arg Thr Leu Ala Pro Gln  
-5 1 5  
atg tgc tca tct ttt gct acg gga ccc aga caa tac gat gga ata ttc 152  
Met Cys Ser Ser Phe Ala Thr Gly Pro Arg Gln Tyr Asp Gly Ile Phe  
10 15 20  
tat gaa ttt cgt tct tat tac ctt aag ccc tca aag atg aat gag ttc 200  
Tyr Glu Phe Arg Ser Tyr Tyr Leu Lys Pro Ser Lys Met Asn Glu Phe  
25 30 35  
ctg gaa aat ttt gag aaa aac gct caa ctt cgg aca gct cac tct gaa 248  
Leu Glu Asn Phe Glu Lys Asn Ala Gln Leu Arg Thr Ala His Ser Glu  
40 45 50 55  
ttg gtt gga tac tgg agt gta kaa ttt gga ggc aga atg awt aca gtg 296  
Leu Val Gly Tyr Trp Ser Val Xaa Phe Gly Gly Arg Met Xaa Thr Val  
60 65 70  
ttt cat att tgg aag tat gat aat ttt gct cat cga act gaa ttt cag 344  
Phe His Ile Trp Lys Tyr Asp Asn Phe Ala His Arg Thr Glu Phe Gln  
75 80 85  
aaa gcc ttg gcc aaa gat aag gaa tgg caa gaa caa ttc ctc att cca 392  
Lys Ala Leu Ala Lys Asp Lys Glu Trp Gln Glu Gln Phe Leu Ile Pro  
90 95 100  
aat ttg gct ctc aat tgataaacia gatagtgaga ttacttatct ggtaccatgg 447

Asn Leu Ala Leu Asn

105

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cctgggtgggc	cagctctgtg	gggtgatgca	tttaaaagg	cagttcatgc	tcagtcaat	567
ctaggctaca	caaaactagt	tggagtgttc	cacacagagt	acggagcact	caacagagtt	627
catgttcttt	ggtggaatga	gagtgcagat	agtcgtgcag	ctgggagaca	taagtcccat	687
gaggatccca	gagttgtggc	agctgttcgg	gaaagtgtca	actacctagt	atctcagcag	747
aatatgcttc	tgattcctac	atcgttttca	ccactgaaat	agttttctac	tgaaatacaa	807
aacatttcat	taactgctat	aggatctgtc	tgctaattgt	gcttaaattc	tccaagagg	867
ttctcacttt	tatttgaagg	aggtggtaag	ttaatttgct	atgtttcttg	cattatgaag	927
gctacatctg	tgctttgtaa	gtaccacttc	aaaaaatakt	tctgtttact	ttctgcatgg	987
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<222> 201..332

<221> sig\_peptide

<222> 201..251

<223> Von Heijne matrix

score 7.80000019073486

seq VLWLISFFTFDTG/HG

<221> polyA\_site

<222> 869..880

<400> 324

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gatactttct	ttccaaacag	cataagaagt	gattgancca	caagtatact	gaaggmargg	120
yhccwsvvar	tyctggwgtg	amgagataaa	tcaccagtca	cagactatgc	accgactgc	180
tgctgttcag	tccagggaaa	atg aaa gtt	gga gtg ctg	tgg ctc att	tct ttc	233
		Met Lys Val Gly Val Leu Trp Leu Ile Ser Phe				
		-15		-10		
ttc acc ttc	act gac ggc	cac ggt ggc	ttc ctg ggg	gtg agt tgg	tgc	281
Phe Thr Phe Thr	Asp Gly His Gly	Gly Phe Leu Gly	Val Ser Trp Cys			
-5	1	5	10			
tat gtc tca	tat ctc ttc	tca act aac	tct cct ctc	tgg ttc	cgg cgc	329
Tyr Val Ser Tyr	Leu Phe Ser Thr	Asn Ser Pro	Leu Ser Phe	Arg Arg		
	15	20	25			
att tagaaccct	cactctctag	gggactgcaa	ctgcataatt	taatgtactt		382
Ile						
gagatcagaa	gtcctgagtt	ctcgtttcaa	cattaccaac	attcactgtg	tggccttgga	442
taagtragtc	atttcacctc	ttcggagctt	agatgatcma	actgcaarag	gaggatcttt	502
gattamacta	tcttagagat	cttttccagt	tcaacacatg	ctgtactatg	gcttctcgga	562
tgcagaaaaa	tcacatggat	ggacattagc	aatccttara	cactgtcttt	cctgtctaca	622
ctcgtttgag	tgatgckttc	atctaggatc	atggttttaa	tattctctac	atgctgatga	682
ctcccagctg	tatagctcca	tctcagaacc	tctcccctgt	ccacactcac	atatccatta	742
cctacgtgtt	atttccagct	gggaaatcca	gcggaacctc	ggnaacttca	tttgnntcaa	802
aatcгнаacc	caatccttct	tgcttatctc	agcaagtggg	atcactatct	ttccagctac	862
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<221> sig\_peptide  
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seq MCLLTALVTQVIS/LR

<221> polyA\_site  
<222> 1206..1217

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graagrwgga aasctgtcta aaaatattca agtgtgcaac caaggattta gatgaagcca 180  
gcaaacaaag gaatcatgta atcaggacct gagcga atg tgc tta ctc acg gcg 234  
Met Cys Leu Leu Thr Ala  
-10  
tta gtt aca cag gtg att tcc tta aga aaa aat gca gag aga act tgt 282  
Leu Val Thr Gln Val Ile Ser Leu Arg Lys Asn Ala Glu Arg Thr Cys  
-5 1 5  
tta tgc aag agg aga tgg ccc tgg ngc ccc tcg ccc cgg atc tac tgc 330  
Leu Cys Lys Arg Arg Trp Pro Trp Xaa Pro Ser Pro Arg Ile Tyr Cys  
10 15 20 25  
tca tcc acc cca tgc gat tcc aaa ttc ccc acc gtc tac tcc agt gcc 378  
Ser Ser Thr Pro Cys Asp Ser Lys Phe Pro Thr Val Tyr Ser Ser Ala  
30 35 40  
cca ttc cat gcc ccc ctc ccc gtc cag aat tcc tta tgg ggg cac ccg 426  
Pro Phe His Ala Pro Leu Pro Val Gln Asn Ser Leu Trp Gly His Pro  
45 50 55  
ctc cat ggt tgt tcc tgg caa tgc cac cat ccc cag gga car aat ctc 474  
Leu His Gly Cys Ser Trp Gln Cys His His Pro Gln Gly Gln Asn Leu  
60 65 70  
cag cct gcc agt ctc cad acc cat ctc tcc aag ccc aag cgc cat ttt 522  
Gln Pro Ala Ser Leu Xaa Thr His Leu Ser Lys Pro Lys Arg His Phe  
75 80 85  
ara aar aar rra tgt caa gcc tgatgaarac atgagtggca aaaacattgc 573  
Xaa Lys Lys Xaa Cys Gln Ala  
90 95  
aatgtacara aatgagggtt tctatgctga tccttacctt tatcacgagg gacggatgag 633  
catascctca tcccatgggtg gacacccact ggatgtcccc gaccacatca ttgcatatca 693  
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gagcgtgca ggattatcca gccttgtaga cctcggccct cctctaattgg agaagcaagt 1053

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cattacaagt tatagcaaar atgctgtctag ctaaaaaaaaa aaaa 1217
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<221> sig\_peptide  
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score 4.09999990463257  
seq GILILWIIRLLFS/KT

<221> polyA\_signal  
<222> 930..935

<221> polyA\_site  
<222> 948..959

<400> 326

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Glu Met Val Gln Ala Leu Tyr Glu Ala Pro Ala Tyr His Leu Ile Leu
-30 -25 -20 -15
gaa ggg att ctg atc ctc tgg ata atc aga ctt ctt ttc tct aag act 146
Glu Gly Ile Leu Ile Leu Trp Ile Ile Arg Leu Leu Phe Ser Lys Thr
-10 -5 1
tac aaa tta caa gaa cga tct gat ctt aca gtc aag gaa aaa gaa gaa 194
Tyr Lys Leu Gln Glu Arg Ser Asp Leu Thr Val Lys Glu Lys Glu Glu
5 10 15
ctg att gaa gag tgg caa cca gaa cct ctt gtt cct cct gtc cca aaa 242
Leu Ile Glu Glu Trp Gln Pro Glu Pro Leu Val Pro Pro Val Pro Lys
20 25 30
gac cat cct gct ctc aac tac aac atc gtt tca ggc cct cca agc cac 290
Asp His Pro Ala Leu Asn Tyr Asn Ile Val Ser Gly Pro Pro Ser His
35 40 45 50
aaa act gtg gtg aat gga aaa gaa tgt ata aac ttc gcc tca ttt aat 338
Lys Thr Val Val Asn Gly Lys Glu Cys Ile Asn Phe Ala Ser Phe Asn
55 60 65
ttt ctt gga ttg ttg gat aac cct agg gtt aag gca gca gct tta gca 386
Phe Leu Gly Leu Leu Asp Asn Pro Arg Val Lys Ala Ala Ala Leu Ala
70 75 80
tct cta aag aag tat ggc gtg ggg act tgt gga ccc tgt gga ttt tat 434
Ser Leu Lys Lys Tyr Gly Val Gly Thr Cys Gly Pro Cys Gly Phe Tyr
85 90 95
ggc aca ttt gaa tgaaratgaa ggatcattga tttccttggtg tatggataat 486
Gly Thr Phe Glu
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100

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ccgggaacag gccaaactaaa tatttgatga atgtatgatt tcaaatacag tgaattccct 546
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cctgaaaatg ttattaaata attgtttagg ccgggcatgg tggctcatgc ctgtaatccc 666
agcactttca aaggctgagg caggcagatc acctgaggtc aggagttcaa aaccagcctg 726
gccaacatgc tgaaacctcg tctctactaa aaatacaaaa attagctggg cgtgggtggtg 786
grtgccctgtg gtcccagctr cgtgggaggg tgaggtggga gaattgcttc aacctgggag 846
gcggagggtt cagtgagccg agatcatgcc actgcactcc agcctgggca acagagcaag 906
actgtctcaa aaataaataa ataaataaaa ttgttttaat gaaaaaaaaa aaa 959

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<220>  
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 <222> 29..724

<221> sig\_peptide  
 <222> 29..118  
 <223> Von Heijne matrix  
 score 3.90000009536743  
 seq VAHALSLPAESYG/NX

<221> polyA\_signal  
 <222> 886..891

<221> polyA\_site  
 <222> 910..920

<400> 327

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Glu Pro Val Ser Gly Glu Leu Val Ser Val Ala His Ala Leu Ser Leu
-20 -15 -10
cca gca gag tcg tat ggy aac grt yct gac att gag atg gct tgg gcc 148
Pro Ala Glu Ser Tyr Gly Asn Xaa Xaa Asp Ile Glu Met Ala Trp Ala
-5 1 5 10
atg aga gca atg cag cat gct gaa gtc tat tac aag ctg att tca tca 196
Met Arg Ala Met Gln His Ala Glu Val Tyr Tyr Lys Leu Ile Ser Ser
15 20 25
gtt gac cca cag ttc ctg aaa ctc acc aaa gta gat gac caa att tac 244
Val Asp Pro Gln Phe Leu Lys Leu Thr Lys Val Asp Asp Gln Ile Tyr
30 35 40
tct gag ttc cgg aaa aat ttt gag acc ctt agg ata gat gtg ttg grc 292
Ser Glu Phe Arg Lys Asn Phe Glu Thr Leu Arg Ile Asp Val Leu Xaa
45 50 55
cca gaa gan ctc aag tca gaa tca gcn aaa gag ccc cca gga tac aat 340
Pro Glu Xaa Leu Lys Ser Glu Ser Ala Lys Glu Pro Pro Gly Tyr Asn
60 65 70
tct ttg cca ttg aaa ttg ctc gga acc ggg aag gct ata aca aag ctg 388
Ser Leu Pro Leu Lys Leu Leu Gly Thr Gly Lys Ala Ile Thr Lys Leu

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75	80	85	90	
ttt ata tca gtg ttc agg aca aag aag gag aga aag gag tca aca atg				436
Phe Ile Ser Val Phe Arg Thr Lys Lys Glu Arg Lys Glu Ser Thr Met				
95	100	105		
gag gag aaa aaa gag ctg aca gtg gag aag aag aga aca cca aga atg				484
Glu Glu Lys Lys Glu Leu Thr Val Glu Lys Lys Arg Thr Pro Arg Met				
110	115	120		
gag gag aga aag gag ctg ata gtg gag aag aaa aag agg aag gaa tca				532
Glu Glu Arg Lys Glu Leu Ile Val Glu Lys Lys Lys Arg Lys Glu Ser				
125	130	135		
aca gag aag aca aaa ctg aca aag gag gag aaa aag gga aag aag ctg				580
Thr Glu Lys Thr Lys Leu Thr Lys Glu Glu Lys Lys Gly Lys Lys Leu				
140	145	150		
aca aag aaa tca aca aaa gtg gtg aaa aag cta tgt aag gta tac agg				628
Thr Lys Lys Ser Thr Lys Val Val Lys Lys Leu Cys Lys Val Tyr Arg				
155	160	165	170	
gaa cag cac tct aga agc tat gac tca att gag act aca agt acc acg				676
Glu Gln His Ser Arg Ser Tyr Asp Ser Ile Glu Thr Thr Ser Thr Thr				
175	180	185		
gtg cta ctt gca cag acc cct ttg gtt aaa tgt aaa ttc ttg tac aat				724
Val Leu Leu Ala Gln Thr Pro Leu Val Lys Cys Lys Phe Leu Tyr Asn				
190	195	200		
tgaaaggatac gcagaaggac atctttctag tctaacagtc aggagctgct ctgggtcattc				784
ccttgatatga actggtctaa agactgttag tggggtgtta gttgattttt cctggtatac				844
tgtttcttgg ctgacactac tgggtcaagta agaaatttgt aaataaattt cttttggttc				904
ttattaamaa aaaaaaas				921

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 score 4.09999990463257  
 seq SLMFFSMMATCTS/NV

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ataattttaat gcaaaatatc cttttatgaa tttcatgtta atattgtgaa atattaaaaat	60
aattccacaa tagttgagaa aaatgagcat ttttttccat ttttaaaaaa tgcatagaaa	120
agacaattttt aaaatcctgg gamccawatt tatttagaag tagctgttag taaaacatta	180
gaaaaggagt caggccatba ggttattttat nbnaatctct aagcaattag gntgaagtta	240
ttaagtcaag cctagaaaag ctgcctcctt gtaaggcttt catgacaatg tatagtaatc	300
brcagtgtcc aattcttcgc actcctcagg aatatcacta cctcaggtta cggtaacacag	360

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gctataattg atgatgatgt tcagataact gaagacacaa taa atg aca ttc aga      415
                                   Met Thr Phe Arg
                                   -20
cat cag gac aat tcc ctc atg ttc ttt tct atg atg gcc acc tgt acc      463
His Gln Asp Asn Ser Leu Met Phe Phe Ser Met Met Ala Thr Cys Thr
      -15                    -10                    -5
agc aac gtg ggt ttc acc cac aca acg atg aac tgt tct ctt act tct      511
Ser Asn Val Gly Phe Thr His Thr Thr Met Asn Cys Ser Leu Thr Ser
      1                    5                    10                    15
cca gtt gat ttt aaa gac ttg tta aga gtc tta cta ata aaa ttt ggg      559
Pro Val Asp Phe Lys Asp Leu Leu Arg Val Leu Leu Ile Lys Phe Gly
      20                    25                    30
tat gat aga aaa tcc aca atc aaa tct tgaaccaaat aacatattaa      606
Tyr Asp Arg Lys Ser Thr Ile Lys Ser
      35                    40
attactaata tttaagtgat ggaagacaca caaaaaactt aaaagcacga acaacctaac      666
ttgaaaaara attttaaaat atgattaacc tgaaraaaaa araatcctaa ragccaaagc      726
tcctttttat ttagcttgga attttccctaa tgggttcctaa caaactgtcc caatgtcata      786
taaggaaaca tgatctatta cattccttta taacaacgtg gararactat aaacctatgt      846
aagtagtaaa actatatcag adactcagga ractgactww aaggcctgga tctgcagtgt      906
attatctgta taaaaattgg cagggggaag ctaaaaggaa aggagattgg agatctcaat      966
tctatcatgg tgtatttcat acgcaaatac ragcatgcat tgttttttgt ttttggaaar      1026
avaarggaag tgtgtttctgc cccatgtttc cttccgtgtt tatagttcaa actctatata      1086
cacttcaggt attttttgtt tagcccttca ttataaatgg gcaggaaatt gtttatcaac      1146
ctagccagtt tattactagt gaccttgact tcagtatcct gagcattcct ttatatTTTT      1206
cttttattat cctgagtcctg taactaaaca attttgtcct caaattttta tccaatatcc      1266
attgcaccac accaaatcaa gcttcttgat tttcaaaaat aaaaaggggg aaatacttac      1326
aacttgtaaa aaaaaaaaaa                                     1344

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 score 7  
 seq AGLSSCLLPLCWL/ER

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aagcctaggt gtggcgcccc gaccggactt tcacttctgg ccagcccttt cccacactgg      60
gcgcggggass ggtgccagtc tttaaacaac ctctcgatgg gtcccacgaa gatgtttcca      120
gacccttgga atgccaaagt caagtttagc tatgtctcgc ggagaggccg gtggaagaag      180
caacgagaat gaagcacccc agttctctgc tgagcacatg ggcatctgca ataaagattt      240

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aatttccag cttctcctga agctcggtat ggccacaaca ctaaattctg cccgaggaga 300
ttgagcaaaa tagtatggga cttccaagaa atg ttt tta aag tca ggg gca ggc 354
                               Met Phe Leu Lys Ser Gly Ala Gly
                               -15
ctt tct tca tgc ctt ctt cct ctt tgc tgg ctg gaa cgc aaa gac cat 402
Leu Ser Ser Cys Leu Leu Pro Leu Cys Trp Leu Glu Arg Lys Asp His
-10 -5 1 5
ggc agg agg cca agc asc cat cct gga agg tgaaagcctc atactaagga 452
Gly Arg Arg Pro Ser Xaa His Pro Gly Arg
10 15
cgtcaracag cgaaataara rcctgggtcc ttgacctgt aaasatctcc ctccccatcc 512
tggtctgtct gccttgactc ctttcatatg aaaaaataaa acttttaact tgcgtwaacc 572
aaaaaaaaaaa aaa 585
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score 5.09999990463257  
seq FLLSQMSQHQVHA/VQ  
  
<221> polyA\_signal  
<222> 886..891  
  
<221> polyA\_site  
<222> 903..914  
  
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acaaatatca atgatgttta tgaatctagt gtgaaagtkt taatcacatc acaaggct 58
atg aac rra tat gca agt cca ttc aac tgw caa ttg ard tat ttg gak 106
Met Asn Xaa Tyr Ala Ser Pro Phe Asn Xaa Gln Leu Xaa Tyr Leu Xaa
-50 -45 -40
ttg agc agr ttc gag tgt gtr cat aga gat gga aga gta att aca ctg 154
Leu Ser Arg Phe Glu Cys Val His Arg Asp Gly Arg Val Ile Thr Leu
-35 -30 -25
tct tat cag gag cag gag cta cag gat ttt ctt ctg tct cag atg tca 202
Ser Tyr Gln Glu Gln Glu Leu Gln Asp Phe Leu Leu Ser Gln Met Ser
-20 -15 -10
cag cac cag gta cat gca gtt cag caa ctc gcc aag gtt atg ggc tgg 250
Gln His Gln Val His Ala Val Gln Gln Leu Ala Lys Val Met Gly Trp
-5 1 5 10
caa gta ctg agc ttc agt aat cat gtg gga ctt gga cct ata gag agc 298
Gln Val Leu Ser Phe Ser Asn His Val Gly Leu Gly Pro Ile Glu Ser
15 20 25
abt ggt aat gca tct gcc atc acg gtg gcc ccc caa gtg gtg act atg 346
Xaa Gly Asn Ala Ser Ala Ile Thr Val Ala Pro Gln Val Val Thr Met
30 35 40
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cta ttt cag ttc gta atg gac ctg aaa gtg gca gca aga tta tgg ttc	394
Leu Phe Gln Phe Val Met Asp Leu Lys Val Ala Ala Arg Leu Trp Phe	
45 50 55	
agt ttc ctc gta acc aat gta aar acc ttc caa aaa gtg atg ttt tac	442
Ser Phe Leu Val Thr Asn Val Lys Thr Phe Gln Lys Val Met Phe Tyr	
60 65 70	
aar ata aca aat gga gtc atc ttc gtg ggc cat tca aar aag ttc agt	490
Lys Ile Thr Asn Gly Val Ile Phe Val Gly His Ser Lys Lys Phe Ser	
75 80 85 90	
gga ata aaa tgg aag gtc kaa att ttg ttt ata aaa tgg arm tgc tta	538
Gly Ile Lys Trp Lys Val Xaa Ile Leu Phe Ile Lys Trp Xaa Cys Leu	
95 100 105	
tgt ctg cac tta gcc ctt gtc tac tat gat ttt ttc car atg ttt cct	586
Cys Leu His Leu Ala Leu Val Tyr Tyr Asp Phe Phe Gln Met Phe Pro	
110 115 120	
aaa raa gtt tcc ara aac ttt gac ttg aaa tgt ttg car atc aac tat	634
Lys Xaa Val Ser Xaa Asn Phe Asp Leu Lys Cys Leu Gln Ile Asn Tyr	
125 130 135	
aag cac aaa gaa gar ata act tcc aaa aga gtg ctg ttt tta aaa ata	682
Lys His Lys Glu Glu Ile Thr Ser Lys Arg Val Leu Phe Leu Lys Ile	
140 145 150	
ata att agg aaa tgt ttt att tagcactttc aaacttttca ctttataaat	733
Ile Ile Arg Lys Cys Phe Ile	
155 160	
gacaagtgtc ttgaaatgca gaagtttatg tacagttgta tatacagtat gacaagatgt	793
aaaataaatat gtttttcatg cagtttaaaa tattactaac ttaagggttt ctatgtgctt	853
ttttaaataat tccttctttg atgttgacat caaataaagt atgtggttta aaaaaaaaaa	913
	914

<210> 331

<211> 1161

<212> DNA

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<221> CDS

<222> 672..752

<221> sig\_peptide

<222> 672..722

<223> Von Heijne matrix

score 4.30000019073486

seq LLYAHLSTSKRA/VV

<221> polyA\_site

<222> 1150..1161

<400> 331

aagatatcac tgtcttgttt tcacttagat cctacttaca aagtgagggt tattaacaga	60
ataaagcctt ccttttaaagc tttataataa tcatatttat taataatgct gttgtgcata	120
cttatagtat gcatatattc agcatatggt gcatgtsttc agaattacat aagatgaaat	180
ccctttcatt gcaacttgca agtgagaaaa gatccttagt ggctctggtg gaagaaatag	240
tattttcttct tctcaggggtg tctccctgcc ttggcccttc ccagaagccc cggctttaaa	300
agtgaaaatg tttgaaacat gaaacatgtc tgtaggaagc atcagcatgg ccataagtgc	360
artgattttc atatatgcct ctgcccattt caaatatatt tttgacatga ataaatctaa	420

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cagtatacar aataattcat gtaaraccct aacgtgtaca tgtgaaaaag catttctata 480
taatgtgagg agcactggcc atcaattagg gaaataaagg tcatgtaata ttgcaaattt 540
tcaaaataga gcsstgcaag ataactgcaa tcataccaaa aactatttga gtaaatggat 600
ttttaagta atttttgttt aaaaaaatat atatttcaga agsagaaaat gtcaaagat 660
agtctttgta a atg gtg gtg cac ctt ctc tat gca cat ctg tct ttt aca 710
          Met Val Val His Leu Leu Tyr Ala His Leu Ser Phe Thr
                -15                -10                -5
tca aaa aga gct gtg gtc atg cta aaa tta gag ata act ttt 752
Ser Lys Arg Ala Val Val Met Leu Lys Leu Glu Ile Thr Phe
                1                5                10
tgaatgactt ggtcaagctg tgtgtaaaat atttaacat aagtcaagta cagtgtacta 812
tgtttaataa agttacattt aatgcattta ttgcatatat gaatatatac atgaagaggc 872
tttatgtctt ctggtatttg attttgaatg ttttttaagt cagtgggtgcc tttaggcaag 932
aactttcgaa attaatacatt ctttgtgttt tctgattttt caggtaacat gtacactatt 992
tagaaacat catagtttat tcaccttaaa aaattgattg tattatttaa atatatcact 1052
tagatgggca tttcctataa ttaggatatt ccaaatagtt gctgaaatca attgtgccat 1112
tgaccaatgg atgcacttgg ttagccttaa ttttttyaaa aaaaaaaaaa 1161
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<212> DNA
<213> Homo sapiens
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<220>
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<222> 57..311
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      score 5.30000019073486
      seq LFHLLFLPHYIET/FK
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<222> 332..337
<221> polyA_site
<222> 351..363
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<400> 332
acattttctta ctgccttacg ctcacacctga ggtccacctt ggtctctaaa aacacc atg 59
                                   Met
tgt tct cat gcc tcc atg tct ttt cac aca ctg ttc cat ttg ctc ttc 107
Cys Ser His Ala Ser Met Ser Phe His Thr Leu Phe His Leu Leu Phe
          -20          -15          -10
ctc cca cat tac att gaa act ttc aag cct cag tcg aaa cat tgc ttc 155
Leu Pro His Tyr Ile Glu Thr Phe Lys Pro Gln Ser Lys His Cys Phe
          -5          1          5
ttc tgg ata gca gcc ttc ttg aca tcc ctc ctc act ccc cag tcc cta 203
Phe Trp Ile Ala Ala Phe Leu Thr Ser Leu Leu Thr Pro Gln Ser Leu
10          15          20          25
cag ggc ttc cat agc tct tta tgt gca ctt cga tcc cag cat ttt cca 251
Gln Gly Phe His Ser Ser Leu Cys Ala Leu Arg Ser Gln His Phe Pro
          30          35          40
tcg act tgt aat tgt ttc tgc tac ctg aca atc atc gcc ttg drd tac 299
```

Ser Thr Cys Asn Cys Phe Cys Tyr Leu Thr Ile Ile Ala Leu Xaa Tyr  
45 50 55  
tgg gac aac ctt tgattactca ttatatcctc aataaatatt tgttgaacca 351  
Trp Asp Asn Leu  
60  
aaaaaaaaaa aa 363

<210> 333  
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<212> DNA  
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<220>  
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<222> 80..232

<221> sig\_peptide  
<222> 80..127  
<223> Von Heijne matrix  
score 3.70000004768372  
seq IALTLIPSMLSR/AG

<221> polyA\_signal  
<222> 617..622

<221> polyA\_site  
<222> 634..645

<400> 333  
accttcttgt tatttatgct attctctttg tggctccatt cttctttcaa tcttctcagc 60  
ttataacogt ctttccctt atg cta agg ata gcc ctt aca ctc atc cca tct 112  
Met Leu Arg Ile Ala Leu Thr Leu Ile Pro Ser  
-15 -10  
atg ctg tca agg gct gct ggt tgg tgc tgg tac aag gag ccc act cag 160  
Met Leu Ser Arg Ala Ala Gly Trp Cys Trp Tyr Lys Glu Pro Thr Gln  
5 1 5 10  
cag ttt tct tac ctt tgc ctg ccc tgc ctt tca tgg aat aar aaa ggc 208  
Gln Phe Ser Tyr Leu Cys Leu Pro Cys Leu Ser Trp Asn Lys Lys Gly  
15 20 25  
aac gtt ttg cag ctt cca aat ttc tgaaraaact aatctcarat tggcagttaa 262  
Asn Val Leu Gln Leu Pro Asn Phe  
30 35  
agtcaaaatg ttgccaaata tttattcctt ttgcctaakt ttggctaccc ggttcaattg 322  
ctttttatatt ttaatgtctt gactcttcar agttcgtacc tcaaaaraac aatgaraaca 382  
tttgctttgc tttctgctga atccctaatac tcaacaatct atacctggac tgtccagttc 442  
tcttctgtg ctatcttctc ttctatccaa gtaraatgta ygccaggarc tcttccctc 502  
tarcaatttc tactaaaatg tccaagtara atgtttcctt ttacaatcaa attactgtat 562  
ttattaattt gctaraatcc aktaaatacat tttggtagct ctggetgtgc tatcaataaa 622  
aagatgaaag caaaaaaaaaaaa aaa 645

<210> 334  
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<213> Homo sapiens

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<221> CDS

<222> 91..291

<221> sig\_peptide

<222> 91..219

<223> Von Heijne matrix

score 3.79999995231628

seq LISVLYLIPKTLT/TN

<221> polyA\_signal

<222> 367..372

<221> polyA\_site

<222> 389..400

<400> 334

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taaaatatatt tgttcttcaa ttacagagcg atg acc cca cag tat ctg cct cac      114
                               Met Thr Pro Gln Tyr Leu Pro His
                               -40
ggg gga aaa tac caa gtt ctt gga gat tac tct ttg gca gtg gtc ttc      162
Gly Gly Lys Tyr Gln Val Leu Gly Asp Tyr Ser Leu Ala Val Val Phe
-35                               -30                               -25                               -20
ccc ctg cac ttt tct gat cta att tct gtt tta tac ctt ata ccc aaa      210
Pro Leu His Phe Ser Asp Leu Ile Ser Val Leu Tyr Leu Ile Pro Lys
                               -15                               -10                               -5
aca ctt act acc aac aca gct gtt aaa cat tct ata caa aaa aat tgt      258
Thr Leu Thr Thr Asn Thr Ala Val Lys His Ser Ile Gln Lys Asn Cys
                               1                               5                               10
atg mat ctg gta tta gga aaa tta ctt tca cag taaatatcaa agaaaaaaga      311
Met Xaa Leu Val Leu Gly Lys Leu Leu Ser Gln
15                               20
ttaagggtct ctttgccatg cttttcatca tatgcaccaa atgtaaattt tgtacaataa      371
taattttattt cctaagyaaa aaaaaaaaaa      400
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<210> 335

<211> 496

<212> DNA

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<221> CDS

<222> 196..384

<221> sig\_peptide

<222> 196..240

<223> Von Heijne matrix

score 6.699999980926514

seq ILSTVTALTFARA/LD

<221> polyA\_signal

<222> 461..466



<221> polyA\_site  
<222> 485..496

<400> 335

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cccgagata ggaccaaccg tcaggaatgc gaggaatgtt tttcttcgga ctctatcgag      180
gcacacagac agacc atg ggg att ctg tct aca gtg aca gcc tta aca ttt      231
          Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe
          -15                -10                -5
gcc aga gcc ctg gac ggc tgc aga aat ggc att gcc cac cct gca agt      279
Ala Arg Ala Leu Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser
          1                5                10
gag aag cac aga ctc gag aaa tgt agg gaa ctc gag agc agc cac tcg      327
Glu Lys His Arg Leu Glu Lys Cys Arg Glu Leu Glu Ser Ser His Ser
          15                20                25
gcc cca gga tca acc cag cac cga aga aaa aca acc aga aga aat tat      375
Ala Pro Gly Ser Thr Gln His Arg Arg Lys Thr Thr Arg Arg Asn Tyr
          30                35                40                45
tct tca gcc tgaaatgaak ccgggatcaa atggttgctg atcaragccc      424
Ser Ser Ala
tatatttaaatt tggaaaagtc aaattgasca ttattaaata aagcttggtt aatatgtctc      484
aaacaaaaaa aa      496
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<210> 336

<211> 968

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 54..590

<221> sig\_peptide

<222> 54..227

<223> Von Heijne matrix

score 3.5

seq GGILMGSFQGTIA/GQ

<221> polyA\_site

<222> 955..965

<400> 336

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atatttgccc cttactttat cttgtgcctt gagaaattgc tggggagaga ggt atg      56
                                     Met
tcc act ggg cag ctg tac agg atg gag gat ata ggg cgt ttc cac tcc      104
Ser Thr Gly Gln Leu Tyr Arg Met Glu Asp Ile Gly Arg Phe His Ser
          -55                -50                -45
cag cag cca ggt tcc ctc acc cca agc tca ccc act gtt ggg gag att      152
Gln Gln Pro Gly Ser Leu Thr Pro Ser Ser Pro Thr Val Gly Glu Ile
          -40                -35                -30
atc tac aat aac acc aga aac aca ttg ggg tgg att ggg ggt atc ctt      200
Ile Tyr Asn Asn Thr Arg Asn Thr Leu Gly Trp Ile Gly Gly Ile Leu
          -25                -20                -15                -10
```

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atg ggt tct ttt cag gga acc att gct gga caa ggc aca gga gcc acc      248
Met Gly Ser Phe Gln Gly Thr Ile Ala Gly Gln Gly Thr Gly Ala Thr
      -5                      1                      5
tcc att tct gag ctc tgc aag gga caa gaa cta gag cca tca ggg gct      296
Ser Ile Ser Glu Leu Cys Lys Gly Gln Glu Leu Glu Pro Ser Gly Ala
      10                      15                      20
ggg ctc act gtg gcc cca ccc caa gcc gtc agc ctc cag gdw atc tac      344
Gly Leu Thr Val Ala Pro Pro Gln Ala Val Ser Leu Gln Gly Ile Tyr
      25                      30                      35
acc ctg cct tgg ctg cta cag ctt ttt cac tcc act gcc cta rgg gna      392
Thr Leu Pro Trp Leu Leu Gln Leu Phe His Ser Thr Ala Leu Xaa Xaa
      40                      45                      50                      55
dtt cag caa cct aat gga tct cta tct ctg aac atc tct tca tcc cat      440
Xaa Gln Gln Pro Asn Gly Ser Leu Ser Leu Asn Ile Ser Ser Ser His
      60                      65                      70
gct ccr rgt cca rca acc tgc acc ctg gaa cca gga gtg gac cct acc      488
Ala Pro Xaa Pro Xaa Thr Cys Thr Leu Glu Pro Gly Val Asp Pro Thr
      75                      80                      85
cga sct gtc tgt att aat ccc cat ccc cca cca cca atc tta aaa abc      536
Arg Xaa Val Cys Ile Asn Pro His Pro Pro Pro Pro Ile Leu Lys Xaa
      90                      95                      100
cct ctg tcc ccc tac cct aaa ccc cag tta ggt acc cat gct ggg caa      584
Pro Leu Ser Pro Tyr Pro Lys Pro Gln Leu Gly Thr His Ala Gly Gln
      105                      110                      115
gtc aat taacaattta tgcacaggta ctagttttat tgtattaccg ttccagggtgta      640
Val Asn
120
gctttgaaaa aagtatctca aaaaggcaac atgggcccag cgcagtggct cacgcctgta      700
atcccgagcac tttgggagggc caaggtgggc agatcgccctg aggtctggag ttcaagacca      760
gcctggccaa caggttgaaa ccccgctctct acaaaaaatar gaaaatttrgc caggtgtggt      820
ggcagacgtc tgtrgtccca gctattcagg agactgaggc acgagaattc catgaaccca      880
ggatgaggag gttgcagtga gccgagattg tgccactgcg ctccagcctg ggcgacagag      940
tggtattctg tttcaaaaaa aaaaamcm
968

<210> 337
<211> 901
<212> DNA
<213> Homo sapiens

<220>
<221> CDS
<222> 133..846

<221> sig_peptide
<222> 133..345
<223> Von Heijne matrix
      score 9.39999961853027
      seq VVSFLLLLAGLIA/TY

<221> polyA_site
<222> 890..901

<400> 337
aagcagcttc caggatcctg agatccggag cagccgggggt cggagcggct cctcaagagt      60
tactgatcta tnnatggcag agaaaaaaaa attgtgacca gagacgtgta gcaatgaaca      120

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aggaacrtca ta atg rwn nnk ttc aca gac ccc tct tca gtg aat gaa aag 171
      Met Xaa Xaa Phe Thr Asp Pro Ser Ser Val Asn Glu Lys
            -70                -65                -60
aag agg agg gag cgg gaa gaa agg cag aat att gtc ctg tgg aga cag 219
Lys Arg Arg Glu Arg Glu Glu Arg Gln Asn Ile Val Leu Trp Arg Gln
            -55                -50                -45
ccg ctc att acc ttg cag tat ttt tct ctg gaa atc ctt gta atc ttg 267
Pro Leu Ile Thr Leu Gln Tyr Phe Ser Leu Glu Ile Leu Val Ile Leu
            -40                -35                -30
aag gaa tgg acc tca aaa tta tgg cat cgt caa agc att gtg gtg tct 315
Lys Glu Trp Thr Ser Lys Leu Trp His Arg Gln Ser Ile Val Val Ser
            -25                -20                -15
ttt tta ctg ctg ctt gct ggg ctt ata gct acg tat tat gtt gaa gga 363
Phe Leu Leu Leu Leu Ala Gly Leu Ile Ala Thr Tyr Tyr Val Glu Gly
            -10                -5                1                5
gtg cat caa cag tat gtg caa cgt ata gag aaa cag ttt ctt ttg tat 411
Val His Gln Gln Tyr Val Gln Arg Ile Glu Lys Gln Phe Leu Leu Tyr
            10                15                20
gcc tac tgg ata ggc tta gga att ttg tct tct gtt ggg ctt gga aca 459
Ala Tyr Trp Ile Gly Leu Gly Ile Leu Ser Ser Val Gly Leu Gly Thr
            25                30                35
ggg ctg cac acc ttt ctg ctt tat ctg ggt cca cat ata gcc tca gtt 507
Gly Leu His Thr Phe Leu Leu Tyr Leu Gly Pro His Ile Ala Ser Val
            40                45                50
aca tta gct gct tat gaa tgc aat tca gtt aat ttt ccc gaa cca ccc 555
Thr Leu Ala Ala Tyr Glu Cys Asn Ser Val Asn Phe Pro Glu Pro Pro
            55                60                65                70
tat cct gat cag att att tgt cca gat gaa gag ggc act gaa gga acc 603
Tyr Pro Asp Gln Ile Ile Cys Pro Asp Glu Glu Gly Thr Glu Gly Thr
            75                80                85
att tct ttg tgg agt atc atc tca aaa gtt agg att gaa gcc tgc atg 651
Ile Ser Leu Trp Ser Ile Ile Ser Lys Val Arg Ile Glu Ala Cys Met
            90                95                100
cgg ggt atc ggt aca gca atc gga gag ctg cct cca tat ttc atg gcc 699
Trp Gly Ile Gly Thr Ala Ile Gly Glu Leu Pro Pro Tyr Phe Met Ala
            105                110                115
aga gca gct cgc ctc tca ggt gct gaa cca gat gat gaa gag tat cag 747
Arg Ala Ala Arg Leu Ser Gly Ala Glu Pro Asp Asp Glu Glu Tyr Gln
            120                125                130
gaa ttt gaa gag atg ctg gaa cat gca gag tct gca caa gta aga aca 795
Glu Phe Glu Glu Met Leu Glu His Ala Glu Ser Ala Gln Val Arg Thr
            135                140                145                150
gtg ggg ata gaa aat aga aca ctt tac ttc ttc cta aag agg cta tta 843
Val Gly Ile Glu Asn Arg Thr Leu Tyr Phe Phe Leu Lys Arg Leu Leu
            155                160                165
agg taaaattggtt agtagttact ctgaagaaga aaactgctaa agtaaaaaaa aaaaa 901
Arg

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<210> 338

<211> 1347

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 138..671

<221> sig\_peptide

<222> 138..248

<223> Von Heijne matrix

score 3.5

seq LVFNFLILITILT/IW

<221> polyA\_signal

<222> 1319..1324

<221> polyA\_site

<222> 1338..1347

<400> 338

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aagaatgctt gtgaagtagc aactaaagtg gcagtgtttc ttctgaaatt ctcaggcagt      60
cagactgtct taggcaaadc ttgataaaat agcccttadc cagggttttta tctaaggaat      120
ccaagaaga ctgggga atg gag aga cag tca agg gtt atg tca gaa aag      170
                Met Glu Arg Gln Ser Arg Val Met Ser Glu Lys
                -35                                -30
gat gag tat cag ttt caa cat cag gga gcg gtg gag ctg ctt gtc ttc      218
Asp Glu Tyr Gln Phe Gln His Gln Gly Ala Val Glu Leu Leu Val Phe
-25                                -20                                -15
aat ttt ttg ctc atc ctt acc att ttg aca atc tgg tta ttt aaa aat      266
Asn Phe Leu Leu Ile Leu Thr Ile Leu Thr Ile Trp Leu Phe Lys Asn
-10                                -5                                1                                5
cat cga ttc cgc ttc ttg cat gaa act gga gga gca atg gtg tat ggc      314
His Arg Phe Arg Phe Leu His Glu Thr Gly Gly Ala Met Val Tyr Gly
10                                15                                20
ctt aya atg gga cta att tta csa tat gct aca gca cca act gat att      362
Leu Xaa Met Gly Leu Ile Leu Xaa Tyr Ala Thr Ala Pro Thr Asp Ile
25                                30                                35
gaa agt ggr rct gtc tat gac tgt gta aaa cta act ttc agt cca tca      410
Glu Ser Gly Xaa Val Tyr Asp Cys Val Lys Leu Thr Phe Ser Pro Ser
40                                45                                50
act ctg ctg gtt aat atc act gac caa gtt tat gar tat aaa tac aar      458
Thr Leu Leu Val Asn Ile Thr Asp Gln Val Tyr Glu Tyr Lys Tyr Lys
55                                60                                65                                70
aga gaa ata agt cag cac amc atc aat cct cat cam gga aat gct ata      506
Arg Glu Ile Ser Gln His Xaa Ile Asn Pro His Xaa Gly Asn Ala Ile
75                                80                                85
ctt gaa aag atg aca ttt gat cca raa atc ttc ttc aat gtt tta ctg      554
Leu Glu Lys Met Thr Phe Asp Pro Xaa Ile Phe Phe Asn Val Leu Leu
90                                95                                100
cca cca att ata ttt cat gca gga tat agt cta aag aag aga cac ttt      602
Pro Pro Ile Ile Phe His Ala Gly Tyr Ser Leu Lys Lys Arg His Phe
105                                110                                115
ttt caa aac tta gga tct att tta acg tat gcc ttc ttg gga act gcc      650
Phe Gln Asn Leu Gly Ser Ile Leu Thr Tyr Ala Phe Leu Gly Thr Ala
120                                125                                130
atc tcc tgc atc gtc ata ggg taagtgcacat tcggagctca agttgcaggt      701
Ile Ser Cys Ile Val Ile Gly
135                                140
ggctgtgggg tcygtgatct gtgtgagggg tctaacactt ccaggattct tgctggckgg      761
gaaaattgtc ttttttttar tawatcacaw atttgtatgt tttttcwgac ttaattccac      821
ggcttckgam aaatacaagg cttcaaataca aagcaaaacta waggattgct ggactttctc      881
tgtgagttct ggactttctga cttaggggaat gtggatcact tgccttgagt tatgtgaagc      941
```

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gcattgcatt cttcttttag tttgagtaat sccgatatgc tcaactgcatt cttttttgtc 1001
ttgtattgag agaccttacc tgtatttggc aggagtgcaa aagtaactat atgccaagag 1061
ttttctttct aaaggaaagt ttacaagaca gcagtctgaa acagatatgt ccaaatatca 1121
acagagttgc ttaatacagg gatagctttt cagttaatac cctgtagaat gcagactctt 1181
tttttcattg tattttcttg attatgctac tgagccctaa gtcacacgtt atatactctg 1241
gcttgcagct catcataaag taaaatgtgg taccaaatgg tgaaggcaat ccagcctctg 1301
ataatcccgt ccaatacatt aaagctccac tgcaggaaaa aaaaaa 1347
```

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<210> 339
<211> 987
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 124..411
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<221> sig_peptide
<222> 124..186
<223> Von Heijne matrix
      score 6.30000019073486
      seq MVALCCCLWKISG/CE
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<221> polyA_signal
<222> 948..953
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```
<221> polyA_site
<222> 971..983
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<400> 339
aagacgctgc ctttagggag agataaaaag cataatgaca ttagctagga aagttaattt 60
tcagttctta ctgaagtgc gtatgaaact gaaatttcca aggaactgaa ttttgtgagc 120
caa atg agc atg caa ttc ttg ttt aag atg gtg gcc tta tgc tgt tgt 168
Met Ser Met Gln Phe Leu Phe Lys Met Val Ala Leu Cys Cys Cys
      -20                      -15                      -10
ctc tgg aag atc tcc ggc tgt gag gaa gtc cct cta act tac aac ctg 216
Leu Trp Lys Ile Ser Gly Cys Glu Glu Val Pro Leu Thr Tyr Asn Leu
      -5                      1                      5                      10
ctc aag tgc ctc cta gat aaa gcg cac tgt gta ctc ctg aca cct tgt 264
Leu Lys Cys Leu Leu Asp Lys Ala His Cys Val Leu Leu Thr Pro Cys
      15                      20                      25
ggg tac atc ttt tcc ttg atc agt cca gaa att ctc aaa ctc act tta 312
Gly Tyr Ile Phe Ser Leu Ile Ser Pro Glu Ile Leu Lys Leu Thr Leu
      30                      35                      40
atc act ttg cav atc ctc tta ata ctc aaa aat cta cac tta ctg tgg 360
Ile Thr Leu Xaa Ile Leu Leu Ile Leu Lys Asn Leu His Leu Leu Trp
      45                      50                      55
ctg aca gtt tca agc awa tgt gtt cat cgc agt agt gca aga aaa gaa 408
Leu Thr Val Ser Ser Xaa Cys Val His Arg Ser Ser Ala Arg Lys Glu
      60                      65                      70
aag tagaagaacc ctgcagagat ttgatggaac ccagcttcta ttcattaataa 461
Lys
75
ccaatggcaa aatataaagc aaataggagg tgacgaaggt tacaaaaata cgtattgttt 521
atgttttccc tgggggtgtgc tgattgtcag gcatcagttc cctgtgccat tcattcccca 581
```

acacagcatg	catcagaaat	tttatcaata	aatgctttct	ctctcaatgt	tcaacctatg	641
ctgatagacc	attaaatata	gtttttgggt	tcacagcttg	tcatcatcat	ttgtctatac	701
ctgtggcaaa	gaatatctaa	taagatactc	tcagcatttt	gcacacttaa	actaagatgc	761
tgaatgctgt	attttacgga	ataatcagcc	acattaaatt	tggagactca	acaagcatgc	821
tgtgaacatt	caacattagg	tttaaatttt	atttttaaaa	gttaataata	aaaggatata	881
tgtaagtat	tatgaaaccc	tgcataact	gtaataaaat	ggtaggatgtg	aatggacaat	941
atatgcaata	aaatttataa	tttgattcya	aaaaaaaaaa	aamccv		987

<210> 340  
<211> 748  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 372..494

<221> sig\_peptide  
<222> 372..443  
<223> Von Heijne matrix  
score 5.30000019073486  
seq RILLLHFYCLLR/SE

<221> polyA\_signal  
<222> 708..713

<221> polyA\_site  
<222> 732..745

<400> 340	
acatgaaatg	tgcttggtct
tgaggttgtg	taattcagct
aaaccgaaggc	agatgagctc
ctttattcct	ttcattgtat
gggtctggtg	ttttgaggta
gaaatgtcta	cagtcagttg
gctttcactt	a atg cac aca ttt gcc
	Met His Thr Phe Ala Asn Asp Arg Gly Leu Tyr Arg Ile
	-20 -15
ctt ctt tta cat ttc tat tgt ctg cta cgc tca tca gag tat att ttg	458
Leu Leu Leu His Phe Tyr Cys Leu Leu Arg Ser Ser Glu Tyr Ile Leu	
-10 -5 1 5	
ggg tac aag gtt ttg ggg gtt ttt tty ccc att ttg taactgcctt	504
Gly Tyr Lys Val Leu Gly Val Phe Phe Pro Ile Leu	
10 15	
attgaaaadt	aaktgccctt
atctggggga	tcatttgtat
tcaattctgt	ggggctcctc
gggatggccc	gacttttgct
accc	
	564
	624
	684
	744
	748

<210> 341  
<211> 1106

<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 112..450

<221> sig\_peptide  
<222> 112..192  
<223> Von Heijne matrix  
score 7.19999980926514  
seq SLLFFLLLEGGXT/EQ

<221> polyA\_signal  
<222> 1053..1058

<221> polyA\_site  
<222> 1095..1106

<400> 341

aagacctcgg aacgagagcg ccccggggag ctcggagcgc gtgcacgcgt ggcavacgga	60
gaaggcvakk rcnnnnrctt gaaggttctg tcaccttttg cagtgggtcca a atg aga	117
	Met Arg
raa aag tgg aaa atg gga ggc atg aaa tac atc ttt tcg ttg ttg ttc	165
Xaa Lys Trp Lys Met Gly Gly Met Lys Tyr Ile Phe Ser Leu Leu Phe	
-25 -20 -15 -10	
ttt ctt ttg cta gaa gga ggc kaa aca gag caa gtr amn cat tca gag	213
Phe Leu Leu Leu Glu Gly Gly Xaa Thr Glu Gln Val Xaa His Ser Glu	
-5 1 5	
aca tat tgc atg ttt caa gac aag aag tac aga gtg ggt gag aga tgg	261
Thr Tyr Cys Met Phe Gln Asp Lys Lys Tyr Arg Val Gly Glu Arg Trp	
10 15 20	
cat cct tac ctg gaa cct tat ggg ttg gtt tac tgc gtg aac tgc atc	309
His Pro Tyr Leu Glu Pro Tyr Gly Leu Val Tyr Cys Val Asn Cys Ile	
25 30 35	
tgc tca gag aat ggg aat gtg ctt tgc agc cga gtc aga tgt cca aat	357
Cys Ser Glu Asn Gly Asn Val Leu Cys Ser Arg Val Arg Cys Pro Asn	
40 45 50 55	
gtt cat tgc ctt tct cct gtg cat att cct cat ctg tgc tgc cct cgc	405
Val His Cys Leu Ser Pro Val His Ile Pro His Leu Cys Cys Pro Arg	
60 65 70	
tgc cca gaa gac tcc tta ccc cca gtg aac aat rwg gtg acc agc	450
Cys Pro Glu Asp Ser Leu Pro Pro Val Asn Asn Xaa Val Thr Ser	
75 80 85	
tagtcttgck agtacaatgg gacaacttac caacatggas agctgttcgt agctgrrggg	510
ctctttcaga atcggaacc cmatcaatgc acccagtgc gctgttcgga rggaaacktg	570
tattgtggtc tcaagacttg ccccaaatta acctgtgcct tcccagtcct tgttccarat	630
tctgtctgcc gggwtgacag argagatgga caactgtcat gggaaacmttc tgatgggtgat	690
atcttcgggc aacctgcca cagagaagca agacattctt accaccgctc tcaactatgat	750
cctccacca gccgacaggc tggaggtctg tcccgctttc ctggggccag aagtcaccgg	810
ggagctctta tggattccca gcaagcatca ggaaccattg tgcaaattgt catcaataac	870
aaacacaagc atggacaagt gtgtgtttcc aatggaaaga cctattotca tggcgagtcc	930
tggcacccaa acctccgggc atttggcatt gtggagtgtg tgctatgtac ttgtaatgtc	990
accaagcaag agtgaagaa aatccactgc cccaatcgat acccctgcaa gtatcctcaa	1050
aaaatagacg gaaaatgctg caaggtgtgt ccaggtaaaaa aagcaaaaaa aaaaaa	1106

<210> 342  
 <211> 1191  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 117..866

<221> sig\_peptide  
 <222> 117..170  
 <223> Von Heijne matrix  
 score 10.6999998092651  
 seq LILLALATGLVGG/ET

<221> polyA\_signal  
 <222> 1159..1164

<221> polyA\_site  
 <222> 1178..1190

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<400> 342
aaaaccagc ctacctgctg tagctgccgc cactgccgtc tccgccgcc ctggwccccc 60
agagcbnmag cccagagacc taggaacctg gggcccgctc ctccccctc caggcc atg 119
Met
agg att ctg cag tta atc ctg ctt gct ctg gca aca ggg ctt gta ggg 167
Arg Ile Leu Gln Leu Ile Leu Leu Ala Leu Ala Thr Gly Leu Val Gly
-15 -10 -5
gga gag acc agg atc atc aag ggg ttc gag tgc aag cct cac tcc cag 215
Gly Glu Thr Arg Ile Ile Lys Gly Phe Glu Cys Lys Pro His Ser Gln
1 5 10 15
ccc tgg cag gca gcc ctg ttc gag aag acg cgg cta ctc tgt ggg gcg 263
Pro Trp Gln Ala Ala Leu Phe Glu Lys Thr Arg Leu Leu Cys Gly Ala
20 25 30
acg ctc atc gcc ccc aga tgg ctc ctg aca gca gcc cac tgc ctc aag 311
Thr Leu Ile Ala Pro Arg Trp Leu Leu Thr Ala Ala His Cys Leu Lys
35 40 45
ccc cgc tac ata ktt cac ctg ggg cag cac aac ctc cag aag gag gag 359
Pro Arg Tyr Ile Xaa His Leu Gly Gln His Asn Leu Gln Lys Glu Glu
50 55 60
ggc tgt gag car acc cgg aca gcc act gag tcc ttc ccc cac ccc ggc 407
Gly Cys Glu Gln Thr Arg Thr Ala Thr Glu Ser Phe Pro His Pro Gly
65 70 75
ttc aac aac agc ctc ccc aac aaa gac cam mgc aat gac atc atg ctg 455
Phe Asn Asn Ser Leu Pro Asn Lys Asp Xaa Xaa Asn Asp Ile Met Leu
80 85 90 95
gtg aak atg gma tcg cca gtc tcc atc acc tgg gct gtg cga ccc ctc 503
Val Xaa Met Xaa Ser Pro Val Ser Ile Thr Trp Ala Val Arg Pro Leu
100 105 110
acc ctc tcc tca cgc tgt gtc act gct ggc acc agc tgc ctc att tcc 551
Thr Leu Ser Ser Arg Cys Val Thr Ala Gly Thr Ser Cys Leu Ile Ser
115 120 125
ggc tgg ggc agc acg tcc agc ccc cag tta cgc ctg cct cac acc ttg 599
Gly Trp Gly Ser Thr Ser Ser Pro Gln Leu Arg Leu Pro His Thr Leu
130 135 140
cga tgc gcc aac atc acc atc att gag cac cag aag tgt gag aac gcc 647
    
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Arg Cys Ala Asn Ile Thr Ile Ile Glu His Gln Lys Cys Glu Asn Ala
 145                      150                      155
tac ccc ggc aac atc aca gac acc atg gtg tgt gcc agc gtg cag gaa      695
Tyr Pro Gly Asn Ile Thr Asp Thr Met Val Cys Ala Ser Val Gln Glu
160                      165                      170                      175
ggg ggc aag gac tcc tgc cag ggt gac tcc ggg ggc cct ctg gtc tgt      743
Gly Gly Lys Asp Ser Cys Gln Gly Asp Ser Gly Gly Pro Leu Val Cys
                      180                      185                      190
aac cag tct ctt caa ggc att atc tcc tgg ggc cag gat ccg tgt gcg      791
Asn Gln Ser Leu Gln Gly Ile Ile Ser Trp Gly Gln Asp Pro Cys Ala
                      195                      200                      205
atc acc cga aag cct ggt gtc tac acg aaa gtc tgc aaa tat gtg gac      839
Ile Thr Arg Lys Pro Gly Val Tyr Thr Lys Val Cys Lys Tyr Val Asp
                      210                      215                      220
tgg atc cag gag acg atg aag aac aat tagactggac ccaccaccca      886
Trp Ile Gln Glu Thr Met Lys Asn Asn
                      225                      230
cagcccatca ccctccattt ccacttggtg tttggttctt gttcactctg ttaataagaa      946
accctaagcc aagaccctct acgaacattc tttgggcctc ctggactaca ggagatgctg      1006
tcacttaata atcaacctgg ggttcgaaat cagtggagacc tggattcaaa ttctgccttg      1066
aaatattgtg actctgggaa tgacaacacc tggtttggtc tctgttgat cccagcccc      1126
aaakwcagct cctggccata tatcaagggt tcaataaata tttgctaaat gaawaaaaaa      1186
aaaaac                                           1191

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<210> 343

<211> 1070

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 13..465

<221> sig\_peptide

<222> 13..75

<223> Von Heijne matrix

score 3.90000009536743

seq PVAVTAAVAPVLS/IN

<221> polyA\_signal

<222> 1035..1040

<221> polyA\_site

<222> 1060..1070

<400> 343

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          Met Ala Ala Ser Thr Ser Met Val Pro Val Ala Val Thr
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gcg gca gtg gcg cct gtc ctg tcc ata aac agc gat ttc tca gat ttg      99
Ala Ala Val Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu
                -5                1                5
cgg gaa att aaa aag caa ctg ctg ctt att gcg ggc ctt acc cgg gag      147
Arg Glu Ile Lys Lys Gln Leu Leu Leu Ile Ala Gly Leu Thr Arg Glu
          10                15                20

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cgg ggc cta cta cac agt agc aaa tgg tgc gcg gag ttg gct ttc tct      195
Arg Gly Leu Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser
25                               30                               35                               40
ctc cct gca ttg cct ctg gcc gag ctg caa ccg cct ccg cct att aca      243
Leu Pro Ala Leu Pro Leu Ala Glu Leu Gln Pro Pro Pro Pro Ile Thr
                               45                               50                               55
gag gaa gat gcc cag gat atg gat gcc tat acc ctg gcc aag gcc tac      291
Glu Glu Asp Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr
                               60                               65                               70
ttt gac gtt aaa gag tat gat cgg gca gca cat ttc ctg cat ggc tgc      339
Phe Asp Val Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys
                               75                               80                               85
aat gca aga aaa gcc tat ttt ctg tat atg tat tcc aga tat ctg gtg      387
Asn Ala Arg Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val
                               90                               95                               100
agg gcc att tta aaa tgt cat tct gcc ttt agt gaa aca tcc ata ttt      435
Arg Ala Ile Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe
105                               110                               115                               120
aga acc aat gga aaa gtt aaa tct ttt aaa tagcttagca gtgggccact      485
Arg Thr Asn Gly Lys Val Lys Ser Phe Lys
                               125                               130
gaatgaatgt actttataca tagcaataat aaaaaaaaga tatcataaat aaagttaaaa      545
aggatggtaa aaaaaaaaaat attcttagga atgactaaca ggataagtaa caacctgatt      605
attttatttac tttaggttat ataaggttct tcatgcctgt gaattaatat tattgtgtaa      665
gaattaagtt aaaaagcctg ggctgacttt taaattttata aattcattta tcatgtttat      725
agtataattta ttgtttttct ttcattggcta ttaaaaagta tgactgtaaa ggacaatgca      785
agtaaaccacaa ctttaactctg tattgaataa taagtacaat ttattatttt actttgaaac      845
tattatgaatt tactttccta ctttttctta gttgttatct atataaattg attaaaaaaa      905
catttttatgt acttttcatt tcctagtaga gggtgagtat cccttatttg aagtgtttgg      965
gacccaaaagt gtttcagatt tcagattttt ttcagatttt ggtatatttg cattatactt     1025
actggttgaa ataaaaaatg ctgcagtgag tgtcaaaaaa aaaaaa                     1070

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<213> Homo sapiens

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<222> 2..718

<221> sig\_peptide  
<222> 2..76  
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score 3.90000009536743  
seq RVGLLLGGGGVYG/SR

<221> polyA\_signal  
<222> 1170..1175

<221> polyA\_site  
<222> 1203..1213

<400> 344  
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Met	Pro	Arg	Lys	Arg	Lys	Cys	Asp	Leu	Arg	Ala	Val	Arg	Val	Gly	Leu	
-25					-20					-15					-10	
tta	ctc	ggt	ggt	ggc	gga	gtc	tac	gga	agc	cgt	ttt	cgc	ttc	act	ttt	97
Leu	Leu	Gly	Gly	Gly	Gly	Val	Tyr	Gly	Ser	Arg	Phe	Arg	Phe	Thr	Phe	
				-5				1				5				
cct	ggc	tgt	aga	gcg	ctt	tcc	ccc	tgg	cgg	gtg	aga	vtg	cag	aga	cga	145
Pro	Gly	Cys	Arg	Ala	Leu	Ser	Pro	Trp	Arg	Val	Arg	Xaa	Gln	Arg	Arg	
		10				15					20					
agg	tgc	gag	atg	agc	act	atg	ttc	gcg	gac	act	ctc	ctc	atc	gtt	ttt	193
Arg	Cys	Glu	Met	Ser	Thr	Met	Phe	Ala	Asp	Thr	Leu	Leu	Ile	Val	Phe	
	25				30					35						
atc	tct	gtg	tgc	acg	gct	ctg	ctc	gca	gag	ggc	ata	acc	tgg	gtc	ctg	241
Ile	Ser	Val	Cys	Thr	Ala	Leu	Leu	Ala	Glu	Gly	Ile	Thr	Trp	Val	Leu	
	40			45					50					55		
gtt	tac	agg	aca	gac	aag	tac	aag	aga	ctg	aag	gca	gaa	gtg	gaa	aaa	289
Val	Tyr	Arg	Thr	Asp	Lys	Tyr	Lys	Arg	Leu	Lys	Ala	Glu	Val	Glu	Lys	
				60				65					70			
cag	agt	aaa	aaa	ttg	gaa	aag	aag	aag	gaa	aca	ata	aca	gag	tca	gct	337
Gln	Ser	Lys	Lys	Leu	Glu	Lys	Lys	Lys	Glu	Thr	Ile	Thr	Glu	Ser	Ala	
		75				80						85				
ggt	cga	caa	cag	aaa	aar	aaa	ata	gag	aga	cdd	kaa	kas	amc	ctg	arg	385
Gly	Arg	Gln	Gln	Lys	Lys	Lys	Ile	Glu	Arg	Xaa	Xaa	Xaa	Xaa	Leu	Xaa	
		90				95						100				
aat	aac	aac	aga	gat	cta	tca	atg	gtt	cga	atg	aaa	tcc	atg	ttt	gct	433
Asn	Asn	Asn	Arg	Asp	Leu	Ser	Met	Val	Arg	Met	Lys	Ser	Met	Phe	Ala	
		105				110					115					
att	ggc	ttt	tgt	ttt	act	gcc	cta	atg	gga	atg	ttc	aat	tcc	ata	ttt	481
Ile	Gly	Phe	Cys	Phe	Thr	Ala	Leu	Met	Gly	Met	Phe	Asn	Ser	Ile	Phe	
	120				125					130				135		
gat	ggt	aga	gtg	gtg	gca	aag	ctt	cct	ttt	acc	cct	ctt	tct	tas	rtc	529
Asp	Gly	Arg	Val	Val	Ala	Lys	Leu	Pro	Phe	Thr	Pro	Leu	Ser	Xaa	Xaa	
				140					145					150		
gga	ctg	tct	cat	cga	aat	ctg	ctg	gga	gat	gac	acc	aca	gac	tgt		577
Xaa	Gly	Leu	Ser	His	Arg	Asn	Leu	Gly	Asp	Asp	Thr	Thr	Asp	Cys		
		155				160						165				
tcc	ttc	att	ttc	ctg	taw	att	ctc	tgt	act	atg	tcg	att	cga	cag	aac	625
Ser	Phe	Ile	Phe	Leu	Xaa	Ile	Leu	Cys	Thr	Met	Ser	Ile	Arg	Gln	Asn	
		170				175						180				
att	cag	aag	att	ctc	ggc	ctt	gcc	cct	tca	cga	gcc	gcc	acc	aag	cag	673
Ile	Gln	Lys	Ile	Leu	Gly	Leu	Ala	Pro	Ser	Arg	Ala	Ala	Thr	Lys	Gln	
		185				190					195					
gca	ggt	gga	ttt	ctt	ggc	cca	cca	cct	cct	tct	ggg	aag	ttc	tct		718
Ala	Gly	Gly	Phe	Leu	Gly	Pro	Pro	Pro	Pro	Ser	Gly	Lys	Phe	Ser		
	200				205					210						
tgaactcaag	aactctttat	tttctakcat	tctttctaga	cacacacaca	tcagactggc											778
aactgttttg	tascaagagc	cataggtagc	cttackactt	gggcctcttt	ctagttttga											838
attattttcta	agccttttgg	gtatkattag	agtgaaaatg	gcagccagca	aacttgatag											898
tgcttttggg	cctagatgat	ttttatcaaa	taagtggatt	gattagttaa	gttcaggtaa											958
tgtttatgta	atgaaaaaca	aatagcatcc	ttcttgtttc	atttacataa	gtattttctg											1018
tgggaccgac	tctcaaggca	ctgtgtatgc	cctgcaagtt	ggctgtctat	gagcatttag											1078
agatttagaa	gaaaaattta	gtttgtttta	cccttgtaac	tgtttgtttt	gttggtgttt											1138
ttttttcaag	ccaaatacat	gacataarat	caataaarag	gccaaatttt	tasctgtttt											1198
atgtaaaaaa	aaaaa															1213

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score 6.30000019073486  
seq LLMSILALIFIMG/NS

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<222> 943..948

<221> polyA\_site  
<222> 963..973

<400> 345

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ggacgaaaga gtcggcgccg ccgta atg cga gag ccg cag aag aga acc gca	112
Met Arg Glu Pro Gln Lys Arg Thr Ala	
-90 -85	
aca atc gca aaa tyc rrg gcs tva gag ggc ctc cga gac ccc tat ggc	160
Thr Ile Ala Lys Xaa Xaa Ala Xaa Glu Gly Leu Arg Asp Pro Tyr Gly	
-80 -75 -70	
cgc ctc tgt ggt agc gag cac ccc cga aga cca cct gag cgg ccc gag	208
Arg Leu Cys Gly Ser Glu His Pro Arg Arg Pro Pro Glu Arg Pro Glu	
-65 -60 -55	
gaa gac ccg agc act cca gag gag gcc tct acc acc cct gaa gaa gcc	256
Glu Asp Pro Ser Thr Pro Glu Ala Ser Thr Thr Pro Glu Glu Ala	
-50 -45 -40	
tcg agc act gcc caa gca caa aag cct tca gtg ccc cgg agc aat ttt	304
Ser Ser Thr Ala Gln Ala Gln Lys Pro Ser Val Pro Arg Ser Asn Phe	
-35 -30 -25 -20	
cag ggc acc aag aaa agt ctc ctg atg tct ata tta gcg ctc atc ttc	352
Gln Gly Thr Lys Lys Ser Leu Leu Met Ser Ile Leu Ala Leu Ile Phe	
-15 -10 -5	
atc atg ggc aac agc gcc aag gaa gct ctg gtc tgg aaa gtg ctg ggg	400
Ile Met Gly Asn Ser Ala Lys Glu Ala Leu Val Trp Lys Val Leu Gly	
1 5 10	
aag tta gga atg cag cct gga cgt cas cac agc atc ttt gga gat ccg	448
Lys Leu Gly Met Gln Pro Gly Arg Xaa His Ser Ile Phe Gly Asp Pro	
15 20 25	
aag aar atc gtc aca gaa ran ttt gtg cgc aga ggg tac ctg att tat	496
Lys Lys Ile Val Thr Glu Xaa Phe Val Arg Arg Gly Tyr Leu Ile Tyr	
30 35 40 45	
ara ccg gtg ccc cgt abc agt ccg gtg gag tat gas ttc ttc tgg ggg	544
Xaa Pro Val Pro Arg Xaa Ser Pro Val Glu Tyr Xaa Phe Phe Trp Gly	
50 55 60	
ccc cga gca cac gtg gaa tcg agc ara ctg aaa stc wtg cat ttt gtg	592
Pro Arg Ala His Val Glu Ser Ser Xaa Leu Lys Xaa Xaa His Phe Val	
65 70 75	
gca agg gtt cgt aac cga tgc tct aaa gac tgg cct tgt aat tat gac	640
Ala Arg Val Arg Asn Arg Cys Ser Lys Asp Trp Pro Cys Asn Tyr Asp	

80	85	90	
tgg gat tgc gac gat gat gca gag gtt gag gct atc ctc aat tca ggt			688
Trp Asp Ser Asp Asp Asp Ala Glu Val Glu Ala Ile Leu Asn Ser Gly			
95	100	105	
gct arg ggt tat tcc gcc cct taagtaratc tgaggcagac ccttggggggt			739
Ala Xaa Gly Tyr Ser Ala Pro			
110	115		
gtaaaagaga gtcacaggta ccccaaggag tagatgccag ggtcctaagt tgaaaatgmt			799
gtcgattggg ggcgggggac actgtatttg atatttgatga tcagtgatca ttgttcaact			859
gcgaaataga gtgtttgctt ttgataatgg aaaattgtat tcgtttttaa attccgtttg			919
ttgagaataa caatatgttt aaaaatataa ttgaacaaat tttaaaaaaa aaaamcccy			978

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 <222> 63..320  
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 <222> 63..179  
 <223> Von Heijne matrix  
 score 3.90000009536743  
 seq VLAIGLLHIVLLS/IP

<221> polyA\_signal  
 <222> 771..776

<221> polyA\_site  
 <222> 799..810

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gg atg aat gtk ggc aca gcg cac ags dag gtg aac ccc aac acg cgg	107
Met Asn Val Gly Thr Ala His Xaa Xaa Val Asn Pro Asn Thr Arg	
-35 -30 -25	
gtk atg aac agc cgt ggc atc tgg ctc tcc tac gtg ctg gcc atc ggt	155
Val Met Asn Ser Arg Gly Ile Trp Leu Ser Tyr Val Leu Ala Ile Gly	
-20 -15 -10	
ctc ctc cac atc gtg ctg ctg agc atc ccg ttt gtk agt gtc cct gtc	203
Leu Leu His Ile Val Leu Leu Ser Ile Pro Phe Val Ser Val Pro Val	
-5 1 5	
gtc tgg acc ctc acc aac ctc att cac aac atg ggc atg tat atc ttc	251
Val Trp Thr Leu Thr Asn Leu Ile His Asn Met Gly Met Tyr Ile Phe	
10 15 20	
ctg cac acg gtg aag ggg aca ccc ttt gag acc ccg gac cag ggc aag	299
Leu His Thr Val Lys Gly Thr Pro Phe Glu Thr Pro Asp Gln Gly Lys	
25 30 35 40	
gcg agg ctg cta acc cac tgg tgagcagatg gattatgggg tccagttcac	350
Ala Arg Leu Leu Thr His Trp	
45	
ggcctctcgg aaktcttga ccatcacacc catcgtgctg tacttctcta ccagcttcta	410
cactaaktac raccaaattc attttgtgct caacaccgtg tccctgatra gcgtgcttat	470

ccccaagctg	ccccagctcc	acggaktccg	gattttttgga	atcaataakt	actgaaaktg	530
casccccctt	ccctgcccag	ggtggcaggg	gaggggtagg	gtaaaaggca	tktgctgcaa	590
chctgaaaac	araaaraara	rsccctctgga	cactgccara	ratggggggtt	gagcctctgg	650
cctaattttcc	cccctcgctt	cccccagtag	ccaacttgga	gtagcttgta	ytgggggttg	710
ggtaggcccc	ctgggctctg	accttttctg	aattttttga	tcttttcctt	ttgctttttg	770
aatararact	ccatggaggt	ggtcatggaa	aaaaaaaaaa			810

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 <222> 299..418

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 <223> Von Heijne matrix  
 score 3.59999990463257  
 seq LTLILLITPSPSPL/LF

<221> polyA\_signal  
 <222> 739..744

<221> polyA\_site  
 <222> 762..771

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taagaaaaga gcaagctgtc cagagagtga gaagtttgaa aagagagggtg cataagagag	120
aatgatgtc catttgagcc ccaccacgga ggttatgtgg tcccaaaaagg aatgatggcc	180
aagcaattaa tttttcctcc tagttcttag cttgcttctg cattgattgg ctttacacaa	240
ctggcattta gtctgcatta cacaaataga cactaattta tttggaacaa gcagcaaa	298
atg aga act tta ttt ggt gca gtc agg gct cca ttt agt tcc ctc act	346
Met Arg Thr Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr	
-25 -20 -15	
ctg ctt cta atc acc cct tct ccc agc cct ctt cta ttt gat aga ggt	394
Leu Leu Leu Ile Thr Pro Ser Pro Ser Pro Leu Leu Phe Asp Arg Gly	
-10 -5 1 5	
ctg tcc ctc aga tca gca atg tct tagccccctct cctctcttcc attccttcc	448
Leu Ser Leu Arg Ser Ala Met Ser	
10	
ggttggtactc atttcttcta actttttaata aacatttagg tataatacat tacagtaagt	508
gctattttaga tacaaactta aaacatacta tatatttttaa ggatctaaga atcctttara	568
rrrggcacat gactgaagta cctcagctgc gcagcctgta accagttttt ttaatgtaaa	628
agtaaraatg ccagccttaa cctabccctg carataaaaag ctaactttta ttaataccag	688
ccctgaataa tggcactaat ccacactctt ccttaragtg atgctggaaa aataaaatca	748
ggggcttcag attaaaaaaa aaa	771

<210> 348  
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 <212> DNA

<213> Homo sapiens

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<221> CDS

<222> 186..380

<221> sig\_peptide

<222> 186..233

<223> Von Heijne matrix

score 4

seq FFLFLSFVLMYDG/LR

<221> polyA\_signal

<222> 383..388

<221> polyA\_site

<222> 396..409

<400> 348

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aattgttagag	aatcattttg	gtgctcaagt	ctcttagcag	tgccattattg	cctcatagca	180
agaag atg	ctg ggg ttt	ttt ttg ttt	ttg tcc ttt	gta tta atg	tat gat	230
Met Leu Gly Phe	Phe Leu Phe	Leu Ser Phe	Val Leu Met	Tyr Asp		
-15	-10	-5				
ggt ttg	cgc ctt	ttt ggc	att ctt	tca aca	tgt cgt	gta cat
Gly Leu Arg	Leu Phe Gly	Ile Leu Ser	Thr Cys Arg	Val His His	Thr	278
1	5	10	15			
atg aat	cag ttc	cta att	gat ata	tct agc	ttt acc	tcc cga
Met Asn Gln	Phe Leu Ile	Asp Ile Ser	Ser Phe Thr	Ser Arg Val	Lys	326
20	25	30				
aaa aaa	atc ttt	tta ttt	tat gcc	ttc awa	ggt tgc	ycg ttt
Lys Lys Ile	Phe Leu Phe	Tyr Ala Phe	Xaa Gly Cys	Xaa Phe	Gln Ser	374
35	40	45				
gcc aca	taaataaaat	gtttaacaaa	aaaaaaaaa			409
Ala Thr						

<210> 349

<211> 613

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 69..458

<221> sig\_peptide

<222> 69..233

<223> Von Heijne matrix

score 4

seq AALCGISLSQLFP/EP

<221> polyA\_signal

<222> 564..569

<221> polyA\_site  
<222> 602..613

<400> 349

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cgctggga atg gcc atg tgg aac agg cca tgb bag ang ctg cct cag cag	110
Met Ala Met Trp Asn Arg Pro Xaa Xaa Xaa Leu Pro Gln Gln	
-55 -50 -45	
cct cts sta gct gag ccc act gca gag ggg gag cca cac ctg ccc acg	158
Pro Leu Xaa Ala Glu Pro Thr Ala Glu Gly Glu Pro His Leu Pro Thr	
-40 -35 -30	
ggc cgg gas byg act gag gcc aac cgc ttc gcc tat gct gcc ctc tgt	206
Gly Arg Xaa Xaa Thr Glu Ala Asn Arg Phe Ala Tyr Ala Ala Leu Cys	
-25 -20 -15 -10	
ggc atc tcc ctg tcc cag tta ttt cct gaa ccc gaa cac agc tcc ttc	254
Gly Ile Ser Leu Ser Gln Leu Phe Pro Glu Pro Glu His Ser Ser Phe	
-5 1 5	
tgc aca gag ttc atg gca ggc ctg gtg ckm tgg ctg gag ttg tct gaa	302
Cys Thr Glu Phe Met Ala Gly Leu Val Xaa Trp Leu Glu Leu Ser Glu	
10 15 20	
gct gtc ttg cca acc atg act gct ttt gcg agc ggc ctg gga ggt gaa	350
Ala Val Leu Pro Thr Met Thr Ala Phe Ala Ser Gly Leu Gly Gly Glu	
25 30 35	
gga sca vma tgt gtt tgt tca aat ttt act gaa gga ccc cat ctt gaa	398
Gly Xaa Xaa Cys Val Cys Ser Asn Phe Thr Glu Gly Pro His Leu Glu	
40 45 50 55	
gga cga ccc gac ggt gat cac tca gga cct tct gag ctt ctc act caa	446
Gly Arg Pro Asp Gly Asp His Ser Gly Pro Ser Glu Leu Leu Thr Gln	
60 65 70	
gga tgg gca cta tgacscgccg gccagagtcc tcgtttgcc catgacctcc	498
Gly Trp Ala Leu	
75	
ctgctccaag tgcccttgga ggagctggat gtccttgaaa agatgttcct ggagagcctg	558
aaggaaatca aagaagagga atctgaaatg gccgaggcat cccraaaaaa aaaaa	613

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<222> 12..638

<221> sig\_peptide  
<222> 12..263  
<223> Von Heijne matrix  
score 4.19999980926514  
seq ITMLQMLALLGYG/LF

<221> polyA\_signal  
<222> 951..956

<221> polyA\_site  
<222> 975..985



<400> 350

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      Met Val Asn Phe Pro Gln Lys Ile Ala Gly Glu Leu Tyr
                                -80                                -75
gga cct ctc atg ctg gtc ttc act ctg gtt gct atc cta ctc cat ggg      98
Gly Pro Leu Met Leu Val Phe Thr Leu Val Ala Ile Leu Leu His Gly
      -70                                -65                                -60
atg aag acg tct gac act att atc cgg gag ggc acc ctg atg ggc aca      146
Met Lys Thr Ser Asp Thr Ile Ile Arg Glu Gly Thr Leu Met Gly Thr
      -55                                -50                                -45                                -40
gcc att ggc acc tgc ttc ggc tac tgg ctg gga gtc tca tcc ttc att      194
Ala Ile Gly Thr Cys Phe Gly Tyr Trp Leu Gly Val Ser Ser Phe Ile
                                -35                                -30                                -25
tac ttc ctt gcc tac ctg tgc aac gcc cag atc acc atg ctg cag atg      242
Tyr Phe Leu Ala Tyr Leu Cys Asn Ala Gln Ile Thr Met Leu Gln Met
                                -20                                -15                                -10
ttg gca ctg ctg ggc tat ggc ctc ttt ggg cat tgc att gtc ctg ttc      290
Leu Ala Leu Leu Gly Tyr Gly Leu Phe Gly His Cys Ile Val Leu Phe
                                -5                                1                                5
atc acc tat aat atc cac ctc cgc gcc ctc ttc tac ctc ttc tgg ctg      338
Ile Thr Tyr Asn Ile His Leu Arg Ala Leu Phe Tyr Leu Phe Trp Leu
      -10                                15                                20                                25
ctg gtg ggt gga ctg tcc aca ctg cgc atg gta gca gtg ttg gtg tct      386
Leu Val Gly Gly Leu Ser Thr Leu Arg Met Val Ala Val Leu Val Ser
                                30                                35                                40
cgg acc gtg ggc ccc aca cad cgg mtg ctc ctc tgt ggc acc ctg gct      434
Arg Thr Val Gly Pro Thr Xaa Arg Xaa Leu Leu Cys Gly Thr Leu Ala
                                45                                50                                55
gcc cta cac atg ctc ttc ctg ctc tat ctg cat ttt gcc tac cac aaa      482
Ala Leu His Met Leu Phe Leu Leu Tyr Leu His Phe Ala Tyr His Lys
                                60                                65                                70
dtg gta dag ggg atc ctg gac aca ctg gag ggc ccc aac atc ccg ccc      530
Xaa Val Xaa Gly Ile Leu Asp Thr Leu Glu Gly Pro Asn Ile Pro Pro
      75                                80                                85
atc cag agg gtc ccc aga gac atc cct gcc atg ctc cct gct gct cgg      578
Ile Gln Arg Val Pro Arg Asp Ile Pro Ala Met Leu Pro Ala Ala Arg
      90                                95                                100                                105
ctt ccc acc acc gtc ctc aac gcc aca gcc aaa gct gtt gcg gtg acc      626
Leu Pro Thr Thr Val Leu Asn Ala Thr Ala Lys Ala Val Ala Val Thr
                                110                                115                                120
ctg cag tca cac tgacccacc tgaaattctt ggccagtcct ctttcccgca      678
Leu Gln Ser His
                                125
gctgcagaga ggargaasac tattaaagga cagtcctgat gacatgtttc gtagatgggg      738
tttgagctg cactgagct gtagctgctg aagtacctcc ttgatgcctg tcggcacttc      798
tgaaaggcac aaggccaaga actcctggcc aggactgcaa ggctctgcag ccaatgcaga      858
aatgggtca gtcctttga gaacccctcc ccacctaccc cttccttctt ctttatctct      918
ccacattgt cttgctaaat atagacttgg taattaaaaat gttgattgaa gtctggaaaa      978
aaaaaaat
                                986

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<210> 351

<211> 1447

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 282..389

<221> sig\_peptide

<222> 282..332

<223> Von Heijne matrix

score 3.5

seq RWWCFHLQAEASA/HP

<221> polyA\_signal

<222> 1413..1418

<221> polyA\_site

<222> 1437..1447

<400> 351

ataataatat	ctaaaaagct	aaatttttaa	taccagcttt	acataaatga	ttgtkgactc	60
tggtctgtkt	ctgacacctt	tccagaaaaa	agtcaattgt	tcaggtacac	caaagaggaa	120
gaagagctgt	ggaggccacc	ctctacaaag	ctttatagaa	cttctggatc	taactcacia	180
acaagcttcc	agaagagact	agagacctta	ggccaggaga	tgaaggagtt	cagtagcaaa	240
gtcacacctg	tccaattccc	tgagctttgc	tcactcagct	a atg gga	tgg caa agg	296

Met Gly Trp Gln Arg

-15

tgg tgg tgc	ttt cat	ctt cag	gca gaa	gcc tct	gcc cat	ccc cct	caa	344
Trp Trp Cys	Phe His	Leu Gln	Ala Glu	Ala Ser	Ala His	Pro Pro	Gln	

-10

-5

1

ggg ctg	cag gcc	caa ttc	tca tgc	tgc cct	tgg gtg	ggc atc	tgt	389
Gly Leu	Gln Ala	Gln Phe	Ser Cys	Cys Pro	Trp Val	Gly Ile	Cys	

5

10

15

taacaaadga	aaacgtctgg	gtggcggcag	casctttgct	ctgagtgcct	acaaagctaa	449
tgcttggtgc	tagaaacatc	atcattatta	aacttcagaa	aagcagcagc	catgttcagt	509
caggctcatg	ctgcctcact	gcttaagtgc	ctgcaggagc	cgcttgccaa	rotccccctc	569
ctacacctgg	cacactgggg	tctgcacaag	gctttgtcaa	ccaaaracag	cttccccccw	629
ttgattgcct	gtagactttg	gagccaaraa	acactctgtg	tgactctaca	cacacttcag	689
gtggtttgtg	cttcaaagtc	attgatgcaa	cttgaaaagga	aacagttaa	tggtggaaat	749
gaactacat	ttataacttc	tgttttttta	ttgagaaaat	gattcacgaa	kkccaaatca	809
gattgccagg	aagaaatagg	acgtgacggg	actgggccct	gtgattctcc	cagcccttgc	869
agtccgctag	gtgagaggaa	aagctcttta	cttccgcccc	tggcagggac	ttctgggtta	929
tgggagaaac	cagagatggg	aatgaggaaa	atatgaaacta	cagcagaagc	ccctgggcag	989
ctgtgatgga	gcccctgaca	ttactcttct	tgcactctgtc	ctgccttctt	tccctctgcg	1049
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cgttggggac	tgccctgtatt	tggaagattt	aaaaacctag	catcctgttc	tcacccctcta	1229
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tgcattcggt	gccaaacattt	tttagagctg	tacccaaaaca	aaaagcctgt	actcacatca	1349
camtgtcatt	ttgataggag	cgtttttgta	tttttacaaag	gcagaatggg	gtgtaacagt	1409
tgaattaaac	ttagcaatca	cgtgctcaaa	aaaaaaaaa			1447

<210> 352

<211> 1641

<212> DNA

<213> Homo sapiens

<220>  
<221> CDS  
<222> 208..339

<221> sig\_peptide  
<222> 208..294  
<223> Von Heijne matrix  
score 5.59999990463257  
seq LFLQLLVSHIVC/AT

<221> polyA\_site  
<222> 1631..1641

<400> 352  
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atgatcatgc agaggcagaa gccaggcacc tgggtctatga atcagaccaa aacaaggatg 120  
gcaagcttac caaggaggag atcgttgaca agtatgactt atttgttggc agccaggcca 180  
cagatttttg ggaggcctta gtacggc atg atg agt tct gag cta cgg agg aac 234  
Met Met Ser Ser Glu Leu Arg Arg Asn

-25  
cct cat ttc ctc aaa agt aat tta ttt tta cag ctt ctg gtt tca cat 282  
Pro His Phe Leu Lys Ser Asn Leu Phe Leu Gln Leu Leu Val Ser His  
-15 -10 -5  
gaa att gtt tgc gct act gag act gtt act aca aac ttt tta aga cat 330  
Glu Ile Val Cys Ala Thr Glu Thr Val Thr Thr Asn Phe Leu Arg His  
1 5 10  
gaa aag gcg taatgaaaac catcccgctc ccattcctcc tctctctctga 379  
Glu Lys Ala

15  
gggactggag ggaagccgtg cttctgagga acaactctaa ttagtacact tgtgtttgta 439  
ratttacacw wtgtattatg tattaacatg gcgtgtttat ttttgtattt ttctctgggt 499  
gggagtatka tatgaaggat caaratcctc aactcacaca tgtaracaaa cattasctct 559  
ttactctttc tcaaccctt wtatgatttt aataattctc acttaactaa ttttgtaagc 619  
ctgagatcaa taagaaatgt tcaggagaga ggaaagaaaa aaaatatatg ctccacaatt 679  
tatatttaga gagagaacac ttagtcttgc ctgtcaaaaa gtccaacatt tcataggtag 739  
tagggggccac atattacatt cagttgctat aggtccagca actgaacctg ccattacctg 799  
ggcaaggaaa gatccctttg ctctaggaaa gcttggccca aattgatttt cttctttttc 859  
ccctgtagg actgactgtt ggctaatttt gtcaagcaca gctgtggtgg gaagagttag 919  
ggccagtgtc ttgaaaatca atcaagtagt gaatgtgatc tctttgcara gctatagata 979  
gaaacagctg gaaaactaaa ggaaaaatac aagtgttttc ggggcataca ttttttttct 1039  
gggtgtgcat ctgttgaaat gctcaagact taattatttg ccttttgaaa tctactgtaaa 1099  
tgcccccatc cggttcctct tcttcccarg tgtgccaagg aattaatctt ggtttcacta 1159  
caattaaaat tcaactcctt ccaatcatgt cattgaaagt gcctttaacg aaagaaatgg 1219  
tcaactgaatg ggaattctct taagaaaccc tgagattaaa aaaagactat ttggataact 1279  
tataggaaaag cctagaacct cccagtagag tggggatttt tttcttcttc cctttctctt 1339  
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tgggctgatt tccaaaaacc acatgctgct gaatttacca gggatcctca tacctcacia 1459  
tgcaaacac ttactaccag gcctttttct gtgtccactg gagagcttga gctcacactc 1519  
aaagatcaga ggacctacag agagggtctt ttggtttgag gaccatggct tacctttcct 1579  
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aa 1641

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<213> Homo sapiens

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<221> CDS

<222> 69..557

<221> sig\_peptide

<222> 69..224

<223> Von Heijne matrix

score 4.69999980926514

seq LGLALGRLEGGSA/RH

<221> polyA\_signal

<222> 849..854

<221> polyA\_site

<222> 870..883

<400> 353

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caagcagg atg gag cac tac cgg aaa gct ggc tct gta gag ctc cca gcg	110
Met Glu His Tyr Arg Lys Ala Gly Ser Val Glu Leu Pro Ala	
-50 -45 -40	
cct tcc cca atg ccc cag cta cct cct gat acc ctt gag atg cgg gtc	158
Pro Ser Pro Met Pro Gln Leu Pro Pro Asp Thr Leu Glu Met Arg Val	
-35 -30 -25	
cga gat ggc agc aaa att cgc aac ctg ctg ggg ttg gct ctg ggt cgg	206
Arg Asp Gly Ser Lys Ile Arg Asn Leu Leu Gly Leu Ala Leu Gly Arg	
-20 -15 -10	
ctg gag ggc ggc agt gct cgg cat gta gtg ttc tca ggt tct ggc agg	254
Leu Glu Gly Gly Ser Ala Arg His Val Val Phe Ser Gly Ser Gly Arg	
-5 1 5 10	
gct gca gga aag gct gtc agc tgc gct gag att gtc aag cgg cgg gtc	302
Ala Ala Gly Lys Ala Val Ser Cys Ala Glu Ile Val Lys Arg Arg Val	
15 20 25	
ccg ggc ctg cac cag ctc acc aag cta ckt ttc ctt caa act gag gac	350
Pro Gly Leu His Gln Leu Thr Lys Leu Xaa Phe Leu Gln Thr Glu Asp	
30 35 40	
agc tgg gtc cca scc tca cct gac aca ggg cta rac ccc ctc aca gtg	398
Ser Trp Val Pro Xaa Ser Pro Asp Thr Gly Leu Xaa Pro Leu Thr Val	
45 50 55	
cgc cgc cat gtg cct gca ktg tgg gtg ctg ctc asc cgg gac ccc ctg	446
Arg Arg His Val Pro Ala Xaa Trp Val Leu Leu Xaa Arg Asp Pro Leu	
60 65 70	
gac ccc aat gag tgt ggt tac caa ccc cca gga gca ccc cct ggc ctg	494
Asp Pro Asn Glu Cys Gly Tyr Gln Pro Pro Gly Ala Pro Pro Gly Leu	
75 80 85 90	
ggt tcc atg ccc agc tcc agc tgt ggc cct cgt tcc cra aaa agg gct	542
Gly Ser Met Pro Ser Ser Ser Cys Gly Pro Arg Ser Xaa Lys Arg Ala	
95 100 105	
cra rac acc cga tcg tgaaaacctg ctgasccagc ctgtttctccg ggcctraatg	597
Xaa Xaa Thr Arg Ser	
110	
tctgggggtgc ttgtgccttt tctranaagc gttgtgaskg ctcaacatcc ccatcaaggt	657
ttgagtcac aaaagtggac ctccctatca tgcttccct tccctctagc atgtgggaag	717
ggactgctgt gaagaatgac agatgtggg cctctgcca gttctgcatt gctaaataag	777
ggcttctct gccttctacc tacagtgcac ttgaactgcc ttctgaaaga ggtccakgga	837

gggatttagg aaataaaagtt tctacctatt tgaaaaaaa aaaacac

884

<210> 354  
<211> 729  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 134..325

<221> sig\_peptide  
<222> 134..274  
<223> Von Heijne matrix  
score 5.90000009536743  
seq TWLGLLSFQNLHC/FP

<221> polyA\_site  
<222> 718..729

<400> 354  
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tgtaatgcaa gtcccctaac tccctgggtg ctaacattaa cttccttaag taataatcaa 120  
tgaaagavat tct atg cat ggt ttt gaa ata ata tcc ttg aaa gag gaa 169  
Met His Gly Phe Glu Ile Ile Ser Leu Lys Glu Glu  
-45 -40  
tca cca tta gga aag gtg agt cag ggt cct ttg ttt aat gtg act agt 217  
Ser Pro Leu Gly Lys Val Ser Gln Gly Pro Leu Phe Asn Val Thr Ser  
-35 -30 -25 -20  
ggc tca tca tca cca gtg acc tgg ttg ggc cta ctc tcc ttc cag aac 265  
Gly Ser Ser Ser Pro Val Thr Trp Leu Gly Leu Leu Ser Phe Gln Asn  
-15 -10 -5  
ctg cat tgc ttc cca gac ctc ccc act gag atg cct cta ara gcc aaa 313  
Leu His Cys Phe Pro Asp Leu Pro Thr Glu Met Pro Leu Xaa Ala Lys  
1 5 10  
gga ktc aac act tgagcctagg gtgggctaca acaaaaratt ctaatttacc 365  
Gly Xaa Asn Thr  
15  
ttgcttcac taggtccagg ccccaakttag cttgctgaag gaacttaaaa agtagctggt 425  
atattattgta ttgtataasc taaaaacatt tatttttgggt gaatcraaac aattccatgt 485  
ascaatcttt tttctgttca cgggtgtttgt gataaaacct taaattccgc aagcatcagt 545  
tttttgaaaa aatgggaatt gaccggatag wwacaggcaa agwtataaat agctacaaca 605  
tcatttaact tttataaaca tgccttctct ctattgaara catctgatat ttttgctgga 665  
aagttggatc tatcctcagt aactctgcca tgaattcctg tttcckgggt ccaaaaaaaa 725  
aaaa 729

<210> 355  
<211> 1013  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS

<222> 78..731

<221> sig\_peptide

<222> 78..227

<223> Von Heijne matrix

score 5.09999990463257

seq RTALILAVCCGSA/SI

<221> polyA\_site

<222> 1002..1013

<400> 355

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aattttatatt acttttct atg cat cat ggc ctc aca cca ctg tta ctt ggt      110
          Met His His Gly Leu Thr Pro Leu Leu Leu Gly
          -50          -45          -40
gta cat gag caa aaa cag caa gtg gtg aaa ttt tta atc aag aaa aaa      158
Val His Glu Gln Lys Gln Gln Val Val Lys Phe Leu Ile Lys Lys Lys
          -35          -30          -25
gca aat tta aat gca ctg gat aga tat gga aga act gct ctc ata ctt      206
Ala Asn Leu Asn Ala Leu Asp Arg Tyr Gly Arg Thr Ala Leu Ile Leu
          -20          -15          -10
gct gta tgt tgt gga tct gca agt ata gtc agc ctt cta ctt gag caa      254
Ala Val Cys Cys Gly Ser Ala Ser Ile Val Ser Leu Leu Leu Glu Gln
          -5          1          5
aac att gat gta tct tct caa gat cta tct gga cag acg gcc aaa aag      302
Asn Ile Asp Val Ser Ser Gln Asp Leu Ser Gly Gln Thr Ala Lys Lys
          10          15          20          25
tat gct gtt tct agt cgt cat aat gta att tgc cag tta ctt tct gac      350
Tyr Ala Val Ser Ser Arg His Asn Val Ile Cys Gln Leu Leu Ser Asp
          30          35          40
tac aaa raa aaa cag atr cta aaa gtc tct tct gaa aac agc aat cca      398
Tyr Lys Xaa Lys Gln Xaa Leu Lys Val Ser Ser Glu Asn Ser Asn Pro
          45          50          55
raa caa gac tta aag ctg aca tca gag gaa gag tca caa agg ctt aaa      446
Xaa Gln Asp Leu Lys Leu Thr Ser Glu Glu Glu Ser Gln Arg Leu Lys
          60          65          70
gga agt gaa aat agc cag cca gag gaa atg tct caa gaa cca gaa ata      494
Gly Ser Glu Asn Ser Gln Pro Glu Glu Met Ser Gln Glu Pro Glu Ile
          75          80          85
aat arg ggt ggt gat aga aag gtt gaa raa raa atg aar aag cac gga      542
Asn Xaa Gly Gly Asp Arg Lys Val Glu Xaa Xaa Met Lys Lys His Gly
          90          95          100          105
agt wct cat atg gga ttc cca raa aac ctg mct aac ggt gcc act gct      590
Ser Xaa His Met Gly Phe Pro Xaa Asn Leu Xaa Asn Gly Ala Thr Ala
          110          115          120
gac aat ggt gat gat gga tta att ccm cca rgg aaa asc ara aca cct      638
Asp Asn Gly Asp Asp Gly Leu Ile Pro Pro Xaa Lys Xaa Xaa Thr Pro
          125          130          135
gaa agc cas caa ttt cct gac act gag aat gaa cag tat cac agg gac      686
Glu Ser Xaa Gln Phe Pro Asp Thr Glu Asn Glu Gln Tyr His Arg Asp
          140          145          150
ttt tct ggc cat ccc mac ttt ccc acd acc ctt ccc atc aaa cag      731
Phe Ser Gly His Pro Xaa Phe Pro Thr Thr Leu Pro Ile Lys Gln
          155          160          165
tgatgaacaa aatgatactc hsaagcmmct ttctgaagam caraacactg gaatattaca      791
agatgagatt ctgattcatg aagaaaagca gatagaagtg gctgaaaatg aattctgagc      851
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tttctcttag ttataaaaaa gaaaaagacc tcttgcatga aaatagtacg ttgcaggaag 911
aaattgtcat gctaaractg gaactagack taatgaaaca tcagagccag ctaaraaaa 971
araaatatth ggaggaaatt gaaagtgtgg aaaaaaaaaa aa 1013
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<210> 356
<211> 973
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 46..693
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<221> sig_peptide
<222> 46..90
<223> Von Heijne matrix
      score 7.59999990463257
      seq CVLVLAAGAVA/VF
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<221> polyA_signal
<222> 937..942
<221> polyA_site
<222> 962..973
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K400> 356
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                                         Met Val Cys Val
                                         -15
ctc gtt cta gct gcg gcc gca gga gct gtg gcg gtt ttc cta atc ctg 105
Leu Val Leu Ala Ala Ala Gly Ala Val Ala Val Phe Leu Ile Leu
-10 -5 1 5
cga ata tgg gta gtg ctt cgt tcc atg gac gtt acg ccc cgg gag tct 153
Arg Ile Trp Val Val Leu Arg Ser Met Asp Val Thr Pro Arg Glu Ser
10 15 20
ctc agt atc ttg gta gtg gct ggg tcc ggt ggg cat acc act gag atc 201
Leu Ser Ile Leu Val Val Ala Gly Ser Gly Gly His Thr Thr Glu Ile
25 30 35
ctg agg ctg ctt ggg agc ttg tcc aat gcc tac tca cct aga cat tat 249
Leu Arg Leu Leu Gly Ser Leu Ser Asn Ala Tyr Ser Pro Arg His Tyr
40 45 50
gtc att gct gac act gat gaa atg agt gcc aat aaa ata aat tct ttt 297
Val Ile Ala Asp Thr Asp Glu Met Ser Ala Asn Lys Ile Asn Ser Phe
55 60 65
gaa cta rat cga gsk gat aga rac cct agt aac atg twt acc aaa tac 345
Glu Leu Xaa Arg Xaa Asp Arg Xaa Pro Ser Asn Met Xaa Thr Lys Tyr
70 75 80 85
tac att cac cga att cca ara agc cgg gag gtt cag cag tcc tgg ccc 393
Tyr Ile His Arg Ile Pro Xaa Ser Arg Glu Val Gln Gln Ser Trp Pro
90 95 100
tcc acc gtt tyc acc acc ttg cac tcc atg tgg ctc tcc ttk ccc cta 441
Ser Thr Val Xaa Thr Thr Leu His Ser Met Trp Leu Ser Xaa Pro Leu
105 110 115
att cac agg gtg aag cca rat ttg gtg ttg tgt aac gga cca gga aca 489
Ile His Arg Val Lys Pro Xaa Leu Val Leu Cys Asn Gly Pro Gly Thr
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120	125	130	
tgt gty cct atc tgt gta tct gcc ctt ctc ctt ggg ata cta gga ata			537
Cys Val Pro Ile Cys Val Ser Ala Leu Leu Leu Gly Ile Leu Gly Ile			
135	140	145	
aag aaa gtg atc att gtc tac gtt gaa agc atc tgc cgt gta aaa acs			585
Lys Lys Val Ile Ile Val Tyr Val Glu Ser Ile Cys Arg Val Lys Thr			
150	155	160	165
tta tcc atg tcc gga aag att ctg ttt cat ctc tca aat tac ttc att			633
Leu Ser Met Ser Gly Lys Ile Leu Phe His Leu Ser Asn Tyr Phe Ile			
170	175	180	
gtt cag tgg ccg gct ctg aaa gaa aag tat ccc aaa tcg gtg tac ctt			681
Val Gln Trp Pro Ala Leu Lys Glu Lys Tyr Pro Lys Ser Val Tyr Leu			
185	190	195	
ggg cga att gtt tgacaaatgg caactgactt ctttagaatt ttgcasttaa			733
Gly Arg Ile Val			
200			
cagtartatg tactcaaatt ggggggaaaa aaaccctaca tgtttcttgt aaaggcgtct			793
gacagtcctg araattattg atggtaagga ataaaaaatg twcagatrac tcagtgaara			853
aactgaggct tctcttatga aacaaacatt gataaacgta actacyaaat gtttatgcct			913
ctgtaaacca aatttctttt ctarataaaa atatgtatta ctacctgcaa aaaaaaaaaa			973

<210> 357  
 <211> 868  
 <212> DNA  
 <213> Homo sapiens

<220>  
 <221> CDS  
 <222> 126..527  
 <221> sig\_peptide  
 <222> 126..182  
 <223> Von Heijne matrix  
 score 3.90000009536743  
 seq ILFHGVFYAGGFA/IV

<221> polyA\_signal  
 <222> 834..839

<221> polyA\_site  
 <222> 856..867

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ggtca atg cct ctg gga gca agg atc ctt ttc cac ggt gtg ttc tat gcc	170
Met Pro Leu Gly Ala Arg Ile Leu Phe His Gly Val Phe Tyr Ala	
-15 -10 -5	
ggg ggc ttt gcc att gtg tat tac ctc att caa aag ttt cat tcc agg	218
Gly Gly Phe Ala Ile Val Tyr Tyr Leu Ile Gln Lys Phe His Ser Arg	
1 5 10	
act tta tat tac aag ttg gca gtg gar cag ctg car arc cat ccc gag	266
Thr Leu Tyr Tyr Lys Leu Ala Val Glu Gln Leu Gln Xaa His Pro Glu	
15 20 25	
gca cag gaa gct ctg ggc cct cct ctc aac atc cat tat ctc aag ctc	314



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Ala Gln Glu Ala Leu Gly Pro Pro Leu Asn Ile His Tyr Leu Lys Leu
  30                      35                      40
atc gac agg gaa aac ttc gtg gac att gtt rat gcc aag ttg aaa att      362
Ile Asp Arg Glu Asn Phe Val Asp Ile Val Xaa Ala Lys Leu Lys Ile
  45                      50                      55                      60
cct gtc tct gga tcc aaa tca gag ggc ctt ctc tac gtc cac tca tcc      410
Pro Val Ser Gly Ser Lys Ser Glu Gly Leu Leu Tyr Val His Ser Ser
                      65                      70                      75
aga ggt ggc ccc ttt cag agg tgg cac ctt gac gag gtc ttt tta gag      458
Arg Gly Gly Pro Phe Gln Arg Trp His Leu Asp Glu Val Phe Leu Glu
                      80                      85                      90
ctc aag gat ggt cag cag att cct gtg ttc aag ctc agt ggg gaa aac      506
Leu Lys Asp Gly Gln Gln Ile Pro Val Phe Lys Leu Ser Gly Glu Asn
                      95                      100                      105
ggt gat gaa gtg aaa aag gag tagagacgac ccagaagacc cagcttgctt      557
Gly Asp Glu Val Lys Lys Glu
  110                      115
ctagtccatc cttccctcat ctctaccata tggccactgg ggtggtggcc catctcagtg      617
acagacactc ctgcaaccca gktttccagc caccagtggg atgatggtat gtgccagcac      677
atggtaattt tgggtgtaatt ctaacttggg cacaacgaat gctatttgtc atttttaaac      737
tgaatccgaa agaaactcct attataaatt taagataatg taatgtattt gaaagtgtt      797
tgataaaaaa agcacatgat aaaaggaatc agaattaata aaatgtttgt tgatctttaa      857
aaaaaaaaaa h                                                                868
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<210> 358

<211> 519

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 66..320

<221> sig\_peptide

<222> 66..113

<223> Von Heijne matrix

score 3.5

seq TALAAXTWLGWVG/VR

<221> polyA\_signal

<222> 490..495

<221> polyA\_site

<222> 508..519

<400> 358

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aattagcgcg taacgcasag actgcttgct gcggcagaga cgccagakgt gcagctccag      60
cagca atg gca gtg acg gcg ttg gcg gcg mrg acg tgg ctt ggc gtg tgg      110
Met Ala Val Thr Ala Leu Ala Ala Xaa Thr Trp Leu Gly Val Trp
  -15                      -10                      -5
ggc gtg agg acc atg caa gcc cga ggc ttc ggc tcg gat cag tcc gag      158
Gly Val Arg Thr Met Gln Ala Arg Gly Phe Gly Ser Asp Gln Ser Glu
  1                      5                      10                      15
aat gtc gac cgg ggc gcg ggc tcc atc cgg gaa gcc ggt ggg gcc ttc      206
Asn Val Asp Arg Gly Ala Gly Ser Ile Arg Glu Ala Gly Gly Ala Phe
```

	20	25	30	
gga aag aga gag cag gct gaa gag gaa cga tat ttc cga gca cag agt				254
Gly Lys Arg Glu Gln Ala Glu Glu Glu Arg Tyr Phe Arg Ala Gln Ser				
	35	40	45	
aca gaa caa ctg gca rct ttg aaa aaa crc cat gaa gaa gar atc gtt				302
Thr Glu Gln Leu Ala Xaa Leu Lys Lys Xaa His Glu Glu Glu Ile Val				
	50	55	60	
cat cat aga gaa gga gat tgagcgtctg cagaaagaaa ttgagcgcca				350
His His Arg Glu Gly Asp				
	65			
taagcagaag atcaaaatgc tagaacatga tgattaagtg cacaccgtgt gccatagaat				410
ggcacatgtc attgccact tctgtgtaaa catgggttctg gtttaactaa tatttgtctg				470
tgtgctacta acagattata ataaattgtc atcagtgaaa aaaaaaaaaa				519

<210> 359  
 <211> 1028  
 <212> DNA  
 <213> Homo sapiens

<220>	
<221> CDS	
<222> 73..948	
<221> sig_peptide	
<222> 73..159	
<223> Von Heijne matrix	
score 4.40000009536743	
seq IVLHLVLQGMVYT/EY	
<221> polyA_site	
<222> 1016..1028	
<400> 359	
agcttttaaag gcctggccag gggaggagca cagatatttt cctgtataat tccagaatgt	60
cttcagagag cc atg cat gga ttg ctt cat tac ctt ttc cat acg aga aac	111
Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn	
	-25 -20
cac acc ttc att gtc ctg cac ctg gtc ttg caa ggg atg gtt tat act	159
His Thr Phe Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr	
	-15 -10 -5
gag tac acc tgg gaa gta ttt ggc tac tgt cag gag ctg gag ttg tcc	207
Glu Tyr Thr Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser	
1 5 10 15	
ttg cat tac ctt ctt ctg ccc tat ctg ctg cta ggt gta aac ctg ttt	255
Leu His Tyr Leu Leu Leu Pro Tyr Leu Leu Leu Gly Val Asn Leu Phe	
	20 25 30
ttt ttc acc ctg act tgt gga acc aat cct ggc att ata aca aaa gca	303
Phe Phe Thr Leu Thr Cys Gly Thr Asn Pro Gly Ile Ile Thr Lys Ala	
	35 40 45
aat gaa tta tta ttt ctt cat gtt tat gaa ttt gat gaa ktg atg ttt	351
Asn Glu Leu Leu Phe Leu His Val Tyr Glu Phe Asp Glu Xaa Met Phe	
	50 55 60
cca aaa aac gtg agg tgc tct act tgt gat tta agg aaa cca gct cga	399
Pro Lys Asn Val Arg Cys Ser Thr Cys Asp Leu Arg Lys Pro Ala Arg	
65 70 75 80	

tcc aas cac tgc akt gtg tgt aac tgg tgt gtg cac cgt ttc rac cat	447
Ser Xaa His Cys Xaa Val Cys Asn Trp Cys Val His Arg Phe Xaa His	
85 90 95	
cac tgt gtt tgg gtg aac aac tgc atc ggg gcc tgg aac atc agg tmc	495
His Cys Val Trp Val Asn Asn Cys Ile Gly Ala Trp Asn Ile Arg Xaa	
100 105 110	
ttc ctc atc tac gtc ttg acc ttg acg gcc tgc gct gcc acc gtc gcc	543
Phe Leu Ile Tyr Val Leu Thr Leu Thr Ala Ser Ala Ala Thr Val Ala	
115 120 125	
att gtg agc acc act ttt ctg gtc cac ttg gtg gtg atg tca gat tta	591
Ile Val Ser Thr Thr Phe Leu Val His Leu Val Val Met Ser Asp Leu	
130 135 140	
tac cag gag act tac atc gat gac ctt gga cac ctc cat gtt atg gac	639
Tyr Gln Glu Thr Tyr Ile Asp Asp Leu Gly His Leu His Val Met Asp	
145 150 155 160	
acg gtc ttt ctt att cag tac ctg ttc ctg act ttt cca cgg att gtc	687
Thr Val Phe Leu Ile Gln Tyr Leu Phe Leu Thr Phe Pro Arg Ile Val	
165 170 175	
ttc atg ctg ggc ttt gtc gtg gtt ctg arc ttc ctc ctg ggt ggc tac	735
Phe Met Leu Gly Phe Val Val Val Leu Xaa Phe Leu Leu Gly Gly Tyr	
180 185 190	
ctg ttg ttt gtc ctg tat ctg gcg gcc acc aac cag act act aac gag	783
Leu Leu Phe Val Leu Tyr Leu Ala Ala Thr Asn Gln Thr Thr Asn Glu	
195 200 205	
tgg tac aga rgt gac tgg gcc tgg tgc cag cgt tgt ccc ctt gtg gcc	831
Trp Tyr Arg Xaa Asp Trp Ala Trp Cys Gln Arg Cys Pro Leu Val Ala	
210 215 220	
tgg cct ccg tca gca gar ccc caa gtc cac cgg aac att cac tcc cat	879
Trp Pro Pro Ser Ala Glu Pro Gln Val His Arg Asn Ile His Ser His	
225 230 235 240	
ggg ctt cgg arc aac ctt caa gar atc ttt cta cct gcc ttt cca tgt	927
Gly Leu Arg Xaa Asn Leu Gln Glu Ile Phe Leu Pro Ala Phe Pro Cys	
245 250 255	
cat gag agg aag aaa caa gaa tgacmagtgt atgactgcct ttgagctgta	978
His Glu Arg Lys Lys Gln Glu	
260	
gtttcccgttt atttacacat gtggatcctc gttttccaaa aaaaaaaaaa	1028

<210> 360

<211> 452

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 69..434

<221> sig\_peptide

<222> 69..236

<223> Von Heijne matrix

score 4.90000009536743

seq FACVPGASPTTLA/FP

<221> polyA\_signal

<222> 419..424

<221> polyA\_site

<222> 441..452

<400> 360

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acagcgtgas tcgcccgccga gaagaatatg aaaaagcaga gcganctcgg ttaagggaaa      60
gcgccgag atg acg ggc ttt ctg ctg ccg ccc gca agc aga ggg act cgg      110
      Met Thr Gly Phe Leu Leu Pro Pro Ala Ser Arg Gly Thr Arg
      -55                    -50                    -45
aga tca tgc agc aga agc aga aaa agg caa acg aga aga agg agg aac      158
Arg Ser Cys Ser Arg Ser Arg Lys Arg Gln Thr Arg Arg Arg Arg Asn
      -40                    -35                    -30
cca agt agc ttt gtg gct tcg tgt cca acc ctc ttg ccc ttc gcc tgt      206
Pro Ser Ser Phe Val Ala Ser Cys Pro Thr Leu Leu Pro Phe Ala Cys
      -25                    -20                    -15
gtg cct gga gcc agt ccc acc acg ctc gcg ttt cct cct gta ktg ctc      254
Val Pro Gly Ala Ser Pro Thr Thr Leu Ala Phe Pro Pro Val Xaa Leu
      -10                    -5                    1                    5
aca ggt ccc avc acc gat ggc att ccc ttt gcc ctr nak tct gca gcg      302
Thr Gly Pro Xaa Thr Asp Gly Ile Pro Phe Ala Leu Xaa Ser Ala Ala
      10                    15                    20
ggt ccc ttt tgt gct tcc ttc ccc tca ggt avc ctc tct ccc cct ggg      350
Gly Pro Phe Cys Ala Ser Phe Pro Ser Gly Xaa Leu Ser Pro Pro Gly
      25                    30                    35
cca ctc ccg ggg gtg agg ggg tta ccc ctt ccc agt gtt ttt tat tcc      398
Pro Leu Pro Gly Val Arg Gly Leu Pro Leu Pro Ser Val Phe Tyr Ser
      40                    45                    50
tgt ggg gct cac ccc aaa gta tta aaa gta gct ttg taattcaaaa      444
Cys Gly Ala His Pro Lys Val Leu Lys Val Ala Leu
      55                    60                    65
aaaaaaaaaaa      452
```

<210> 361

<211> 875

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 628..804

<221> sig\_peptide

<222> 628..711

<223> Von Heijne matrix

score 4.19999980926514

seq LMPVIPALQEAXA/GG

<221> polyA\_site

<222> 864..875

<400> 361

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aaagatggac accgcggagg aagacatatg tagagtgtgt cggtcagaag gaacacctga      60
gaaaccgott tatcatcctt gtgtatgtac tggcagtatt aagttingtcc atcaagaatg      120
cttagttcaa tggctgaaac acagtcgaaa agaatactgt gaattatgca agcacagatt      180
tgcttttaca ccaatttatt ctccagatat gccttcacgg cttccaattc aagacatatt      240
```

tgctggactg gttacaagta ttggcactgc aatacgatat tggtttcatt atacacttgt 300  
 ggccctttgca tggttgggag ttgttctct tacagcatgt gagtattcat gcctctgatt 360  
 ggagttatatt aaacattgca taactactta atattataaa gcaatattgc atcatattat 420  
 tatttgactg atgttttagtt atttgatgtc agagtgtcat gtattaggaa agccttactt 480  
 araaratgtt catcggaact aaraatgakt ttaacaggtc agttttttga gtgaatgtgg 540  
 gaaaraacac agcatacaga atggctaacc atgaaagttc atgaaagcgt kgaaaaaatc 600  
 aaatcaaadc ataattagat atgaagt atg cta rag ctt tca agg gct aca aaa 654

Met Leu Xaa Leu Ser Arg Ala Thr Lys  
 -25 -20

rac ggc cgg gcg cgg tgg ctt atg cct gta atc cca gca ctt cag gag 702  
 Xaa Gly Arg Ala Arg Trp Leu Met Pro Val Ile Pro Ala Leu Gln Glu  
 -15 -10 -5

gcc gan gca ggc gga tca cga ggt cag gag ttt gaa act agc ctg gcc 750  
 Ala Xaa Ala Gly Gly Ser Arg Gly Gln Glu Phe Glu Thr Ser Leu Ala  
 1 5 10

aac atg gag act gag gca gga gaa ttg ctt aaa ccc agg agg cgg agg 798  
 Asn Met Glu Thr Glu Ala Gly Glu Leu Leu Lys Pro Arg Arg Arg Arg  
 15 20 25

ttg car tgaactgaga tcgcaccact gcactccagc ttgggcaaca gagcaagact 854  
 Leu Gln  
 30

ttgtctcgca aaaaaaaaaa a 875

<210> 362

<211> 531

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 70..366

<221> sig\_peptide

<222> 70..108

<223> Von Heijne matrix

score 3.5

seq MHLLSNWANPASS/RR

<221> polyA\_signal

<222> 496..501

<221> polyA\_site

<222> 521..531

<400> 362

aagtggccat ggcggataca gcgactacag catcggcggc ggcggctagt gccgctagcg 60  
 cctcgagcg atg cac ctc ctt tcc aac tgg gca aac ccc gct tcc agc aga 111  
 Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg

-10 -5 1

cgt cct tct atg gcc gct tca ggc act tct tgg ata tca tcg acc ctc 159  
 Arg Pro Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu  
 5 10 15

gca cac tct ttg tca ctg aga gac gtc tca gag agg ctg tgc agc tgc 207  
 Ala His Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys  
 20 25 30

```

tgg agg act ata agc atg gga ccc tgc gcc cgg ggg tca cca atg aac      255
Trp Arg Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn
   35                40                45
agc tct gga gtg cac aga aaa tca agc agg cta ttc tac atc cgg aca      303
Ser Ser Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr
   50                55                60                65
cca atg aga aga tct tca tgc cat tta gaa tgt crg gtt ata ttc ctt      351
Pro Met Arg Arg Ser Ser Cys His Leu Glu Cys Xaa Val Ile Phe Leu
               70                75                80
ttg gga cgc caa ttg taaktgttac cttcaaagga tttccttttc taaaaaatta      406
Leu Gly Arg Gln Leu
               85
ttttaratgt ctaactttat gttattgctc acgggtatatt gactgaattg ttgatttagg      466
ataagtcaat tcctggagggg aaattaccaa ataaaatgat atgtattttct taccacaaaa      526
aaaaa                                                                531

```

<210> 363

<211> 1244

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 70..366

<221> sig\_peptide

<222> 70..108

<223> Von Heijne matrix

score 3.5

seq MHLLSNWANPASS/RR

<221> polyA\_site

<222> 1233..1244

<400> 363

```

tagtggccat ggcggataca gcgactacag catcggcggc ggcggttagt gccgctagcg      60
cctcgagcgc atg cac ctc ctt tcc aac tgg gca aac ccc gct tcc agc aga      111
      Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg
               -10                -5                1
cgt cct tct atg gcc gct tca ggc act tct tgg ata tca tcg acc ctc      159
Arg Pro Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu
               5                10                15
gca cac tct ttg tca ctg aga gac gtc tca gag agg ctg tgc agc tgc      207
Ala His Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys
               20                25                30
tgg agg act ata agc atg gga ccc tgc gcc cgg ggg tca cca atg aac      255
Trp Arg Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn
   35                40                45
agc tct gga gtg cac aga aaa tca agc agg cta ttc tac atc cgg aca      303
Ser Ser Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr
   50                55                60                65
cca atg aga aga tct tca tgc cat tta raa tgt cag gtt ata ttc ctt      351
Pro Met Arg Arg Ser Ser Cys His Leu Xaa Cys Gln Val Ile Phe Leu
               70                75                80
ttg gga cgc caa ttg tagtcggtct tctcttgccc aaccagacac tggcatccac      406

```

Leu Gly Arg Gln Leu

85

tgtcttctg	cagtggctga	accagagcca	caatgcctgt	gtcaactatg	caaaccgcaa	466
tgcraccaag	ccttcacctg	catccaagtt	catccagga	tacctgggag	ctgtcatcag	526
cgccgtctcc	attgctgtgg	gccttatktc	ctggttcaga	aagccaacaa	gttcacccca	586
gccaccgcc	ttctcatcca	gaggtttgtg	ccgttccttg	ctgtagccag	tgccaatatc	646
tgcaatgtgg	tcctgatgcg	gtacggggag	ctggaggaag	ggattgatgt	cctggacagc	706
gatggcaacc	tcgtgggctc	ctccaagatc	gcagcccgc	acgccctgct	ggagacggcg	766
ctgacgcgag	tggtcctgcc	catgcccata	ctgggtgctac	ccccgatcgt	catgtccatg	826
ctggagaaga	cggctctcct	gcaggcacgc	ccccggctgc	tcctccctgt	gcaaagcctc	886
gtgtgcctgg	cagccttcgg	cctggccctg	ccgctggcca	tcagcctctt	cccgcaaagt	946
tcagagattg	aaacatccca	attagagccg	gagatagccc	aggccacgag	cagccggaca	1006
gtggtgtaca	acaaggggtt	gtgagtgtgg	tcagcggcct	ggggacggag	caactgtgcag	1066
ccggggagct	gaggggcarg	gccgtagact	cacggctgca	cctgcaggga	gcagcacgcc	1126
aaccccagca	gtcctgggcc	ccctgggaga	gtgctcaacc	tacagtggag	ggagactgac	1186
ccattcacat	tttaacatag	gcaagaggag	ttctaacaca	tttcgtacaa	aaaaaaaa	1244

<210> 364

<211> 631

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 111..434

<221> sig\_peptide

<222> 111..185

<223> Von Heijne matrix

score 3.90000009536743

seq WIAAVTIAAGTAA/IG

<221> polyA\_site

<222> 618..631

<400> 364

aatcgccgag	tcgggtgcttt	agtagccgc	tggcaccttt	actctcgccg	gccgcgcgaa	60
cccgtttgag	ctcggtatcc	tagtgcacac	gccttgcaag	cgacggcgcc	atg agt	116
					Met Ser	
					-25	
ctg act tcc agt tcc agc gta cga gtt gaa tgg atc gca gca gtt acc						164
Leu Thr Ser Ser Ser Ser Val Arg Val Glu Trp Ile Ala Ala Val Thr						
	-20		-15		-10	
att gct gct ggg aca gct gca att ggt tat cta gct tac aaa aga ttt						212
Ile Ala Ala Gly Thr Ala Ala Ile Gly Tyr Leu Ala Tyr Lys Arg Phe						
	-5		1		5	
tat gtt aaa gat cat cga aat aaa gct atg ata aac ctt cac atc cag						260
Tyr Val Lys Asp His Arg Asn Lys Ala Met Ile Asn Leu His Ile Gln						
10		15		20	25	
aaa gac aac ccc aag ata gta cat gct ttt gac atg gag gat ttg gga						308
Lys Asp Asn Pro Lys Ile Val His Ala Phe Asp Met Glu Asp Leu Gly						
	30		35		40	
gat aaa gct gtg tac tgc cgt tgt tgg agg tcc aaa aag ttc cca ttc						356
Asp Lys Ala Val Tyr Cys Arg Cys Trp Arg Ser Lys Lys Phe Pro Phe						
	45		50		55	

```
tgt gat ggg gct cac aca aaa cat aac gaa gag act gga gac aat gtg      404
Cys Asp Gly Ala His Thr Lys His Asn Glu Glu Thr Gly Asp Asn Val
      60              65              70
ggc cct ctg atc atc aag aaa aaa gaa act taaatggaca cttttgatgc      454
Gly Pro Leu Ile Ile Lys Lys Lys Glu Thr
      75              80
tgcaaatacag cttgtcgtga agttaccta ttgtttaatt araatagacta ccacctctgt      514
ctgattcacc ttcgctggat tctaaatgtg gtatattgcm aactgcagct ttcacattta      574
tggcatttgt cttgttgaaa catcgtggtg cacatttgtt taaacaaaaa aaaaaaa      631
```

<210> 365  
<211> 781  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 19..567

<221> sig\_peptide  
<222> 19..63  
<223> Von Heijne matrix  
score 8.39999961853027  
seq AMWLLCVALAVLA/WG

<221> polyA\_signal  
<222> 749..754  
  
<221> polyA\_site  
<222> 771..781

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<400> 365
aagtgtgtgct tacccatc atg gaa gca atg tgg ctc ctg tgt gtg gcg ttg      51
      Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu
      -15              -10              -5
gcg gtc ttg gca tgg ggc ttc ctc tgg gtt tgg gac tcc tca gaa cga      99
Ala Val Leu Ala Trp Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg
      1              5              10
atg aag agt cgg gag cag gga aga cgg ctg gga gcc gaa agc cgg acc      147
Met Lys Ser Arg Glu Gln Gly Arg Arg Leu Gly Ala Glu Ser Arg Thr
      15              20              25
ctg ctg gtc ata gcg cac cct gac gat gaa gcc atg ttt ttt gct ccc      195
Leu Leu Val Ile Ala His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro
      30              35              40
aca gtg cta ggc ttg gcc cgc cta agg cac tgg gtg tac ctg ctt tgc      243
Thr Val Leu Gly Leu Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys
      45              50              55              60
ttc tct gca gga aat tac tac aat caa gga gag act cgt aag aaa gaa      291
Phe Ser Ala Gly Asn Tyr Tyr Asn Gln Gly Glu Thr Arg Lys Lys Glu
      65              70              75
ctt ttg car agc tgt gat gtt ttg ggg att cca ctc tcc agt gta atg      339
Leu Leu Gln Ser Cys Asp Val Leu Gly Ile Pro Leu Ser Ser Val Met
      80              85              90
att att gac aac agg gat ttc cca rat gac cca ggc atg cag tgg gac      387
Ile Ile Asp Asn Arg Asp Phe Pro Xaa Asp Pro Gly Met Gln Trp Asp
```



```

          95              100              105
aca rag cac gtg gcc ara gtc ctc ctt cag cac ata gaa gtg aat ggc      435
Thr Xaa His Val Ala Xaa Val Leu Leu Gln His Ile Glu Val Asn Gly
      110              115              120
atc aat ctg gtg gtg act ttc gat gca ggg gga rta agt ggc cac agc      483
Ile Asn Leu Val Val Thr Phe Asp Ala Gly Gly Xaa Ser Gly His Ser
      125              130              135              140
aat cac att gct ctg tat gca gct gtg agg aag ctt gag ggc caa att      531
Asn His Ile Ala Leu Tyr Ala Ala Val Arg Lys Leu Glu Gly Gln Ile
              145              150              155
tgc aag ccc tgt ggc act gga caa gac ttt aag gaa tgagtgtgt      577
Cys Lys Pro Cys Gly Thr Gly Gln Asp Phe Lys Glu
              160              165
caatcagtgt gctccacct tcaccatctt cttccctta ctctcacttc cgtcatgtgt      637
tttatacaac tctcaaactt ttcttgaggaga aggaggatat acatacataa tatgaaatgt      697
gtttgttctt cacagtcacc cgattttact gatattttatt tgcattttac caataaaaag      757
aaaatgcaag ctcaaaaaaa aaaa      781
```

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<210> 366
<211> 931
<212> DNA
<213> Homo sapiens
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<220>
<221> CDS
<222> 19..312
<221> sig_peptide
<222> 19..63
<223> Von Heijne matrix
score 8.39999961853027
seq AMWLLCVALAVLA/WG
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<221> polyA_signal
<222> 896..901
<221> polyA_site
<222> 921..931
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```

<400> 366
aagtgtgtgct taccocatc atg gaa gca atg tgg ctc ctg tgt gtg gcg ttg      51
                        Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu
                        -15                        -10                        -5
gcg gtc ttg gca tgg ggc ttc ctc tgg gtt tgg gac tcc tca gaa cga      99
Ala Val Leu Ala Trp Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg
              1              5              10
atg aag agt cgg gag cag gga rga cgg ctg gga gcc gaa agc cgg acc      147
Met Lys Ser Arg Glu Gln Gly Xaa Arg Leu Gly Ala Glu Ser Arg Thr
              15              20              25
ctg ctg gtc ata gcg cac cct gac gat gaa gcc atg ttt ttt gct ccc      195
Leu Leu Val Ile Ala His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro
              30              35              40
aca gtg cta ggc ttg gcc cgc cta agg cac tgg gtg tac ctg ctt tgc      243
Thr Val Leu Gly Leu Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys
45              50              55              60
```

```

ttc tct gca gtt ttc cgt agg gag cta agt gaa tac acc gaa rgt ctt      291
Phe Ser Ala Val Phe Arg Arg Glu Leu Ser Glu Tyr Thr Glu Xaa Leu
      65              70              75
acc tct gaa ccc ctc ama gcc tagggacagg arcggccggc ttacctggtg      342
Thr Ser Glu Pro Leu Xaa Ala
      80
ggttggggga cgtcggcagc tcrctacta cgccagcagg attganganc acagaaacag      402
ttgchsttgg ttgtattcag tacctkcatt tccgttggga actccaccwg tacttggttat      462
kctgtggaac ttttttttat ttgtagaagg agcaagaata ttgaccttac tatatagcac      522
acgaaacaat ctatgctgta tcgtgcctgc tcaatcctta aagttaactt ctaatgatag      582
taaaaracct toctgctgcc tttaaaatgc agcttgtgct aktaacatgc atgtgtcaaa      642
ttgaaraatt agacatagat gactaratar aaagtaattt tgtaggtaat tttaragttc      702
aactccaccc agctttcakt gaaggaacct ttcaaataat aratttttgc ttaccatara      762
raaaaratca aatgacaaag caaatattga ccattaagct ggaatatggt gataattgaa      822
cagttgtata aatgaaktaa ttgaattgta cacatacaat ggggtgaattt tatggcatgt      882
caaagtatac ctcaataaag ctattttttt aaattgcmay aaaaaaaaaa      931

```

<210> 367

<211> 849

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 64..612

<221> sig\_peptide

<222> 64..234

<223> Von Heijne matrix

score 3.79999995231628

seq QLWLVMFCGAGS/VT

<221> polyA\_site

<222> 839..849

<400> 367

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acatacgggc aagtttataa gggctgcat gtcaaacgg gccagcttgc agccatcaag      60
ggt atg gat gtc aca ggg gat gaa gag gaa gaa atc aaa caa gaa att      108
Met Asp Val Thr Gly Asp Glu Glu Glu Glu Ile Lys Gln Glu Ile
      -55              -50              -45
aac atg ttg aag aaa tat tct cat cac cgg aat att gct aca tac tat      156
Asn Met Leu Lys Lys Tyr Ser His His Arg Asn Ile Ala Thr Tyr Tyr
      -40              -35              -30
ggg gct ttt atc aaa aag aac cca cca ggc atg gat gac caa ctt tgg      204
Gly Ala Phe Ile Lys Lys Asn Pro Pro Gly Met Asp Asp Gln Leu Trp
      -25              -20              -15
ttg gtg atg gag ttt tgt ggt gct ggc tct gtc acc gac ctg atc aag      252
Leu Val Met Glu Phe Cys Gly Ala Gly Ser Val Thr Asp Leu Ile Lys
      -10              -5              1              5
aac aca aaa ggt aac acg ttg aaa gag gag tgg att gca tac atc tgc      300
Asn Thr Lys Gly Asn Thr Leu Lys Glu Glu Trp Ile Ala Tyr Ile Cys
      10              15              20
msg gaa atc tta cgg ggg ctg art cac ctg cac cag cat aaa gtg att      348
Xaa Glu Ile Leu Arg Gly Leu Xaa His Leu His Gln His Lys Val Ile
      25              30              35

```

```

cat cga rat att aaa ggg caa aat gtc ttg ctg act gaa aat gca gaa      396
His Arg Xaa Ile Lys Gly Gln Asn Val Leu Leu Thr Glu Asn Ala Glu
  40                      45                      50
gtt aaa cta gtg gac ttt gga rtc akt gct cag ctt gat cga aca gtg      444
Val Lys Leu Val Asp Phe Gly Xaa Xaa Ala Gln Leu Asp Arg Thr Val
  55                      60                      65                      70
ggc agg arg aat act ttc att gga act ccc tac tgg atg gca cca raa      492
Gly Arg Xaa Asn Thr Phe Ile Gly Thr Pro Tyr Trp Met Ala Pro Xaa
                      75                      80                      85
gtt att gcc tgt gat gaa aac cca sat gcc aca tat gat ttc aar art      540
Val Ile Ala Cys Asp Glu Asn Pro Xaa Ala Thr Tyr Asp Phe Lys Xaa
                      90                      95                      100
gac ttg tgg tct ttg ggt atc acc gcc att gaa atg gca gaa ggg ctc      588
Asp Leu Trp Ser Leu Gly Ile Thr Ala Ile Glu Met Ala Glu Gly Leu
                      105                      110                      115
ccc ctc tct gtg aca tgc acc cca tgagagctct cttcctcatc ccccggaatc      642
Pro Leu Ser Val Thr Cys Thr Pro
                      120                      125
cagcgcctcg gctgaagtct aagaagtggc caaaaaaatt ccagtcattt attgagagct      702
gcttggtgtaaa aaatcacagc cagcgaccag caacagaaca attgatgaag catccattta      762
tacgagacca acctaataag cgacaggtcc gcattcaact caaggacatc attgatagaa      822
caagaagaa gcgaggaaaa aaaaaaa      849

```

```

<210> 368
<211> 644
<212> DNA
<213> Homo sapiens

```

```

<220>
<221> CDS
<222> 39..458
<221> sig_peptide
<222> 39..80
<223> Von Heijne matrix
      score 4.40000009536743
      seq FLTALLWRGRIPG/RQ

```

```

<221> polyA_signal
<222> 613..618

```

```

<221> polyA_site
<222> 633..644

```

```

<400> 368
agcggagacg cagagtcttg agcagcgcn caggcacc atg ttc ctg act gcg ctc      56
                                Met Phe Leu Thr Ala Leu
                                -10
ctc tgg cgc ggc cgc att ccc ggc cgt cag tgg atc ggg aag cac cgg      104
Leu Trp Arg Gly Arg Ile Pro Gly Arg Gln Trp Ile Gly Lys His Arg
                      -5                      1                      5
cgg ccg cgg ttc gtg tgc ttg cgc gcc aag cag aac atg atc cgc cgc      152
Arg Pro Arg Phe Val Ser Leu Arg Ala Lys Gln Asn Met Ile Arg Arg
                      10                      15                      20
ctg gag atc gag gcg gag aac cat tac tgg ctg agc atg ccc tac atg      200

```

```

Leu Glu Ile Glu Ala Glu Asn His Tyr Trp Leu Ser Met Pro Tyr Met
25          30          35          40
acc cgg gag cag gag cgc ggc cac gcc gcg ttg cgc agg agg gag gcc      248
Thr Arg Glu Gln Glu Arg Gly His Ala Ala Leu Arg Arg Arg Glu Ala
          45          50          55
ttc gag gcc ata aag gcg gcc gcc act tcc aag ttc ccc ccg cat aga      296
Phe Glu Ala Ile Lys Ala Ala Ala Thr Ser Lys Phe Pro Pro His Arg
          60          65          70
ttc att gcg gac cag ctc gac cat ctc aat vgt cac caa gaa atg gtc      344
Phe Ile Ala Asp Gln Leu Asp His Leu Asn Xaa His Gln Glu Met Val
          75          80          85
cta atc ctg agt cgt cac cct tgg att tta tgg atc acg gag ctg acc      392
Leu Ile Leu Ser Arg His Pro Trp Ile Leu Trp Ile Thr Glu Leu Thr
          90          95          100
atc ttt acc tgg tct gga ctg aaa aac tgt agc ttg tgt gaa aat gag      440
Ile Phe Thr Trp Ser Gly Leu Lys Asn Cys Ser Leu Cys Glu Asn Glu
          105          110          115          120
ctt tgg acc agt ctt tat taaaacaaac aaacatgagt agtctgcata      488
Leu Trp Thr Ser Leu Tyr
          125
tcgaatatct agagctctaa acccccctaat acttaaaagt ctaattgctg tcctgtgggtt      548
tcattagtct gataggaaga tagggatttc ctcagtcaca gatgatattt tgaaggaaag      608
ctgcaataaa gccacaatga tttgaaaaaa aaaaaa      644

```

```

^210> 369
^211> 918
^212> DNA
^213> Homo sapiens

```

```

^220>
^221> CDS
^222> 9..185
^221> sig_peptide
^222> 9..50
^223> Von Heijne matrix
      score 3.70000004768372
      seq AALVTVLFTGVRR/LH

```

```

<221> polyA_site
<222> 906..918

```

```

<400> 369
agctcagc atg gct gct tta gtg act gtt ctc ttc aca ggt gtc cgg agg      50
      Met Ala Ala Leu Val Thr Val Leu Phe Thr Gly Val Arg Arg
          -10          -5
ctg cac tgc agc gcr scg ctt ggg cgg gcg gcc agt ggc grc tac agc      98
Leu His Cys Ser Ala Xaa Leu Gly Arg Ala Ala Ser Gly Xaa Tyr Ser
1          5          10          15
agg aac tgg ctg cca acc cct ccg gct acg ggc ccc tta ccg agc tcc      146
Arg Asn Trp Leu Pro Thr Pro Pro Ala Thr Gly Pro Leu Pro Ser Ser
          20          25          30
cag act ggt cat atg cgg atg gcc gcc ctg ctc ccc caa tgaaaggcca      195
Gln Thr Gly His Met Arg Met Ala Ala Leu Leu Pro Gln
          35          40          45

```

```
gcttcgaaaa aaagctgaaa gggagacktt tgcaaracra kttgtactgc tgtcacagga 255
aatggacgct ggattacaas catggcasct caggcagcar aakttgcagg aaraacaaaag 315
gaagcaggaa aatgctctta aacccaaagg ggcttcactg aaaascccac ttccaaktca 375
ataaaaaagca actcctgcct cccttctca ccctgtctct ggatttcttt tctatcacct 435
aratgcttca tccagccara aaatagcctt cackktcccc atctgtcttc aragcaaaaar 495
agctgggacm ccaaraacaa gctgttarat cactgcctgg gaggcttggc ttartactct 555
catctctggt tccattccag ttcagctaag tcttgcttta aaatttttac ctcttagctg 615
ggtgcggtgg ctacgcctg taatcccagc actttgggag gctgaggcgg gcagatcaca 675
agatcaggag ttcgagacca gcctggccaa cccagcctgg tcaacatggt gaaaccctgt 735
ccctactaaa gatacaaaca attagccggg cgtgggtggg tgcgcttgta atcccagcta 795
ctcaggaggg tgaggcagga gaatcgctta aactcgggag gtagagggtg cagtgagcca 855
aggtcacacc attgcactcc aacctgggag acagggcgag actctgtctc aaaaaaaaaa 915
aaa 918
```

<210> 370  
<211> 472  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 14..316  
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<221> sig\_peptide  
<222> 14..121  
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<223> Von Heijne matrix  
score 5.19999980926514  
seq PLRLNLLILIEG/SV

<221> polyA\_signal  
<222> 442..447  
-----  
<221> polyA\_site  
<222> 458..471  
-----

<400> 370  
attatataga gcc atg ggg cct tac aac gtg gca gtg cct tca gat gta 49  
Met Gly Pro Tyr Asn Val Ala Val Pro Ser Asp Val  
-35 -30 -25  
tct cat gcc cgc ttt tat ttc tta ttt cat cga cca tta agg ctg tta 97  
Ser His Ala Arg Phe Tyr Phe Leu Phe His Arg Pro Leu Arg Leu Leu  
-20 -15 -10  
aat ctg ctc atc ctt att gag ggc agt gtc gtc ttc tat cag ctc tat 145  
Asn Leu Leu Ile Leu Ile Glu Gly Ser Val Val Phe Tyr Gln Leu Tyr  
-5 1 5  
tcc ttg ctg cgg tcg gag aag tgg aac cac aca ctt tcc atg gct ctc 193  
Ser Leu Leu Arg Ser Glu Lys Trp Asn His Thr Leu Ser Met Ala Leu  
10 15 20  
atc ctc ttc tgc aac tac tat gtt tta ttt aaa ctt ctc cgg gac aga 241  
Ile Leu Phe Cys Asn Tyr Tyr Val Leu Phe Lys Leu Leu Arg Asp Arg  
25 30 35 40  
wta kta tta ggc agg gca tac tcc tac cca ctc aac agt tat gaa ctc 289  
Xaa Xaa Leu Gly Arg Ala Tyr Ser Tyr Pro Leu Asn Ser Tyr Glu Leu  
45 50 55  
aag gca aac twa gct gcc tct caw caa tgaggggagaa ctgagataaa 336

Lys Ala Asn Xaa Ala Ala Ser Xaa Gln

60

65

aatatatttca	tacgtttctat	ttttttcttg	tgatattttat	aatatatttaa	gatatttttat	396
atattgtata	ctattatggt	ttgaaagtcg	ggaagagtaa	gggatattaa	atgtatccgt	456
aaacaaaaaa	aaaaam					472

<210> 371

<211> 1504

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 70..1092

<221> sig\_peptide

<222> 70..234

<223> Von Heijne matrix

score 4.09999990463257

seq AVCAALLASHPTA/EV

<221> polyA\_signal

<222> 1475..1480

<221> polyA\_site

<222> 1493..1504

<400> 371

agaaatcgta	ggacttccga	aagcagcggc	ggcgtttgct	tcactgcttg	gaagtgtgag	60
tgcgcggaag	atg cga aag	gtg gtt ttr	att acc ggg	gct agc agt	ggc att	111
	Met Arg Lys	Val Val Leu	Ile Thr Gly	Ala Ser Ser	Gly Ile	
	-55	-50		-45		
ggc ctg gcc	ctc tgc aag	cgg ctg	ctg gcg gaa	gat gat	gag ctt cat	159
Gly Leu Ala	Leu Cys Lys	Arg Leu	Leu Ala Glu	Asp Asp	Glu Leu His	
	-40	-35		-30		
ctg tgt ttg	gcg tgc agg	aat atg	agc aag gca	gaa gct	gtc tgt gct	207
Leu Cys Leu	Ala Cys Arg	Asn Met	Ser Lys Ala	Glu Ala	Val Cys Ala	
	-25	-20		-15	-10	
gct ctg ctg	gcc tct cac	ccc act	gct gag gtc	acc att	gtc cag gtg	255
Ala Leu Leu	Ala Ser His	Pro Thr	Ala Glu	Val Thr	Ile Val Gln	
	-5		1		5	
gat gtc agc	aac ctg cag	tca ttc	ttc cgg gcc	tcc aag	gaa ctt aag	303
Asp Val Ser	Asn Leu Gln	Ser Phe	Phe Arg	Ala Ser	Lys Glu Leu	
	10		15		20	
caa agg ttt	cag aga tta	gac tgt	ata tat cta	aat gct	ggg atc atg	351
Gln Arg Phe	Gln Arg Leu	Asp Cys	Ile Tyr	Leu Asn	Ala Gly Ile	
	25	30		35		
cct aat cca	caa cta aat	atc aaa	gca ctt	ttc ttt	ggc ctc ttt	399
Pro Asn Pro	Gln Leu Asn	Ile Lys	Ala Leu	Phe Phe	Gly Leu Phe	
	40	45		50	55	
aga aaa gtg	att cat atg	ttc tcc	aca gct	gaa ggc	ctg ctg acc	447
Arg Lys Val	Ile His Met	Phe Ser	Thr Ala	Glu Gly	Leu Leu Thr	
	60		65		70	
ggt gat aag	atc act gct	gat gga	ctt cag	gag gtg	ttt gag acc	495
Gly Asp Lys	Ile Thr Ala	Asp Gly	Leu Gln	Glu Val	Phe Glu Thr	
					Asn	

	75		80		85	
gtc ttt ggc cat ttt atc ctg att cgg gaa ctg gag cct ctc ctc tgt						543
Val Phe Gly His Phe Ile Leu Ile Arg Glu Leu Glu Pro Leu Leu Cys						
	90		95		100	
cac agt gac aat cca tct cag ctc atc tgg aca tca tct cgc agt gca						591
His Ser Asp Asn Pro Ser Gln Leu Ile Trp Thr Ser Ser Arg Ser Ala						
	105		110		115	
agg aaa tct aat ttc agc ctc gag gac ttc cag cac agc aaa ggc aag						639
Arg Lys Ser Asn Phe Ser Leu Glu Asp Phe Gln His Ser Lys Gly Lys						
	120		125		130	135
gaa ccc tac agc tct tcc aaa tat gcc act gac ctt ttg agt gtg gct						687
Glu Pro Tyr Ser Ser Ser Lys Tyr Ala Thr Asp Leu Leu Ser Val Ala						
	140		145		150	
ttg aac agg aac ttc aac cag cag ggt ctc tat tcc aat gtg gcc tgt						735
Leu Asn Arg Asn Phe Asn Gln Gln Gly Leu Tyr Ser Asn Val Ala Cys						
	155		160		165	
cca ggt aca gca ttg acc aat ttg aca tat gga att ctg cct ccg ttt						783
Pro Gly Thr Ala Leu Thr Asn Leu Thr Tyr Gly Ile Leu Pro Pro Phe						
	170		175		180	
ata tgg acg ctg ttg atg ccg gca ata ttg cta ctt cgc ttt ttt gca						831
Ile Trp Thr Leu Leu Met Pro Ala Ile Leu Leu Leu Arg Phe Phe Ala						
	185		190		195	
aat gca ttc act ttg aca cca tat aat gga aca gaa gct ctg gta tgg						879
Asn Ala Phe Thr Leu Thr Pro Tyr Asn Gly Thr Glu Ala Leu Val Trp						
	200		205		210	215
ctt ttc cac caa aag cct gaa tct ctc aat cct ctg atc aaa tat ctg						927
Leu Phe His Gln Lys Pro Glu Ser Leu Asn Pro Leu Ile Lys Tyr Leu						
	220		225		230	
agt gcc acc act ggc ttt gga aga aat tac att atg acc cag aag atg						975
Ser Ala Thr Thr Gly Phe Gly Arg Asn Tyr Ile Met Thr Gln Lys Met						
	235		240		245	
gac cta gat gaa gac act gct gaa aaa ttt tat caa aag tta ctg gaa						1023
Asp Leu Asp Glu Asp Thr Ala Glu Lys Phe Tyr Gln Lys Leu Leu Glu						
	250		255		260	
ctg gaa aag cac att agg gtc act att caa aaa aca gat aat cag gcc						1071
Leu Glu Lys His Ile Arg Val Thr Ile Gln Lys Thr Asp Asn Gln Ala						
	265		270		275	
agg ctc agt ggc tca tgc cta taattccagc actttgggag gccaaaggcag						1122
Arg Leu Ser Gly Ser Cys Leu						
	280		285			
aaggatcact tgagaccagg agttcaagac cagcctgaga aacatagtga gcccttgtct						1182
ctacaaaaag aaataaaaat aatagctggg tgtggtggca tgcgcatgta gtcccagcta						1242
ctcagaagga tgaggtggga ggatctcttg aggctgggag gcagaggttg cagtgaactg						1302
agattgtgcc actgcactcc agcctgggtg acagcgagac cctgtctcaa aatatgtata						1362
tatttaatat atatataaaa ccagagctga caatgacact ctggaacatt gcataccttc						1422
tgtacattct ggggtacatg gatttctact gagttggata atatgcattt gtaataaact						1482
atgaactatg aaaaaaaaaa aa						1504

<210> 372

<211> 765

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 274..597

<221> sig\_peptide

<222> 274..399

<223> Von Heijne matrix

score 5.19999980926514

seq LLFDLVCHEFCQS/DD

<221> polyA\_signal

<222> 731..736

<221> polyA\_site

<222> 754..765

<400> 372

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accaggaaca tccagctatt tatgatagca tttgcttcat tatgtcaagt tcaacaaatg      60
ttgacttgct ggtgaagggtg ggggaggttg tggacaagct ctttgatttg gatgagaaac      120
taatgttaag aatgggtcag aaatggggct gctcagcctc tggaccaacc ccaggaagag      180
tctgaagagc agccagtgtt tcggcttggt ccctgtatac ttgaagctgc caaacaagta      240
cgttctgaaa atccagaatg gcttgatggt tac atg cac att tta caa ctg ctt      294
```

Met His Ile Leu Gln Leu Leu

-40

```
act aca gtg gat gat gga att caa gca att gta cat tgt cct gac act      342
Thr Thr Val Asp Asp Gly Ile Gln Ala Ile Val His Cys Pro Asp Thr
35 -30 -25 -20
```

```
gga aaa gac att tgg aat tta ctt ttt gac ctg gtc tgc cat gaa ttc      390
Gly Lys Asp Ile Trp Asn Leu Leu Phe Asp Leu Val Cys His Glu Phe
-15 -10 -5
```

```
tgc cag tct gat gat cca gcc atc att ctt caa raa car aaa acr gtg      438
Cys Gln Ser Asp Asp Pro Ala Ile Ile Leu Gln Xaa Gln Lys Thr Val
1 5 10
```

```
cta gcc tct gtt ttt tca gtg ttg tct gcc atc tat gcc tca cag act      486
Leu Ala Ser Val Phe Ser Val Leu Ser Ala Ile Tyr Ala Ser Gln Thr
15 20 25
```

```
gag caa gak tat cta aar ata raa aaa gga gac ggt ggc tca ggg agt      534
Glu Gln Xaa Tyr Leu Lys Ile Xaa Lys Gly Asp Gly Gly Ser Gly Ser
30 35 40 45
```

```
aaa gga agg cca ktt gan caa aca gaa ktg ttc ctc tgc att tca aaa      582
Lys Gly Arg Pro Xaa Xaa Gln Thr Glu Xaa Phe Leu Cys Ile Ser Lys
50 55 60
```

```
cct tct tcc ttt cta tagccctgtg gtggaagatt ttattaaaat cctacgtgaa      637
Pro Ser Ser Phe Leu
65
```

```
gttgataagg cgcttgctga tgacttgga aaaaacttcc caagtttgaa gggtcagact      697
taaaacctga attggaatta cttctgtaca agaaataaac tttatttttc tcaactgacaa      757
aaaaaaaaa      765
```

<210> 373

<211> 1041

<212> DNA

<213> Homo sapiens

<220>

<221> CDS

<222> 230..469



<221> sig\_peptide  
<222> 230..307  
<223> Von Heijne matrix  
score 4.90000009536743  
seq VLCTNQVLITARA/VP

<221> polyA\_signal  
<222> 1004..1009

<221> polyA\_site  
<222> 1027..1040

<400> 373  
aacttccaag ttgtagtggt gttgtttttca gcctgctgct gctgctgcta ttgcggctag 60  
gggaaccgtc gtggggaagg atggtgtgctg aaaaatgtga aaagaaactt ggtactgtta 120  
tactccaga tacatggaaa gatggtgcta ggaataccac agaaagtggg ggaagaaagc 180  
tgaatgaaaa taaagctttg acttcaaaaa aagccagaat tgatccata atg gaa gaa 238  
Met Glu Glu  
-25  
ata agt tct cca ctt gta gaa ttt gta aaa gtt ttg tgc acc aac cag 286  
Ile Ser Ser Pro Leu Val Glu Phe Val Lys Val Leu Cys Thr Asn Gln  
-20 -15 -10  
gtt ctc att act gcc agg gct gtg cct aca aaa aag gca tct gtg cga 334  
Val Leu Ile Thr Ala Arg Ala Val Pro Thr Lys Lys Ala Ser Val Arg  
-5 1 5  
tgt gtg gaa aaa agg ttt tgg ata cca aaa act aca agc aaa cat ctg 382  
Cys Val Glu Lys Arg Phe Trp Ile Pro Lys Thr Thr Ser Lys His Leu  
10 15 20 25  
tct aga tgt att gat gga att tct ggc ttt cta aat gat ttt act ttc 430  
Ser Arg Cys Ile Asp Gly Ile Ser Gly Phe Leu Asn Asp Phe Thr Phe  
30 35 40  
tgc ctt gaa ttt tca agg cat aga tgt caa ctt aca gaa taacatgkt 479  
Cys Leu Glu Phe Ser Arg His Arg Cys Gln Leu Thr Glu  
45 50  
taagataatt aagtktaaac cagaraattt gattgttact cattttgctc tcatgktcta 539  
aaacagcaac agtghtaacta gtcttttggt gttaatgggt attttcctta taaaaatttt 599  
aaaaactaag tggcaaattc catgaaaata tttctcagtt ctgtatgcac ttttatttaa 659  
cattattcat ataattctcc cccaccact ttatttataa atactgcaaa aktgaraagg 719  
agataataaa tactttgctc tgaatttggc atccaaagtt aacattttctc ccctcactcc 779  
cttgctgggtg tcatagttat tagaatcagc agcctcttaa ctaattgcgg tttcatagga 839  
tatataaatg tttcaagcca ttattgctga atggttcttt agttattaac ctagacccaa 899  
atcaaagacc agttggattt atgatatttt ttatttggtc ttgcagccaa agtgccagtt 959  
tctttaatat gtgaccaaga acacaaggag catccatag gcaaataaa tacactgaat 1019  
tttagaaaaa caaaaaaaaa ar 1041

<210> 374  
<211> 1164  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 72..545

<221> sig\_peptide

<222> 72..203

<223> Von Heijne matrix

score 5.5

seq ILFFTGWIMIDA/AV

<221> polyA\_site

<222> 1151..1162

<400> 374

```

aaagtcggcg tggacgtttg aggaagctgg gatacagcat ttaatgaaaa atttatgctt    60
aagaagtaaa a atg gca ggc ttc cta gat aat ttt cgt tgg cca gaa tgt    110
      Met Ala Gly Phe Leu Asp Asn Phe Arg Trp Pro Glu Cys
                    -40                    -35
gaa tgt att gac tgg agt gag aga aga aat gct gtg gca tct gtt gtc    158
Glu Cys Ile Asp Trp Ser Glu Arg Arg Asn Ala Val Ala Ser Val Val
      -30                    -25                    -20
gca ggt ata ttg ttt ttt aca ggc tgg tgg ata atg att gat gca gct    206
Ala Gly Ile Leu Phe Phe Thr Gly Trp Trp Ile Met Ile Asp Ala Ala
      -15                    -10                    -5                    1
gtg gtg tat cct aag cca gaa cag ttg aac cat gcc ttt cac aca tgt    254
Val Val Tyr Pro Lys Pro Glu Gln Leu Asn His Ala Phe His Thr Cys
                    5                    10                    15
ggg gta ttt tcc aca ttg gct ttc ttc atg ata aat gct gta tcc aat    302
Gly Val Phe Ser Thr Leu Ala Phe Phe Met Ile Asn Ala Val Ser Asn
                    20                    25                    30
gct cag gtg aga ggt gat agc tat gaa agc ggc tgt tta gga aga aca    350
Ala Gln Val Arg Gly Asp Ser Tyr Glu Ser Gly Cys Leu Gly Arg Thr
                    35                    40                    45
ggg gct cga gtt tgg ctt ttc att ggt ttc atg ttg atg ttt ggg tca    398
Gly Ala Arg Val Trp Leu Phe Ile Gly Phe Met Leu Met Phe Gly Ser
                    50                    55                    60                    65
ctt att gct tcc atg tgg att ctt ttt ggt gca tat gtt acc caa aat    446
Leu Ile Ala Ser Met Trp Ile Leu Phe Gly Ala Tyr Val Thr Gln Asn
                    70                    75                    80
act gat gtt tat ccg gga cta gct gtg ttt ttt caa aat gca ctt ata    494
Thr Asp Val Tyr Pro Gly Leu Ala Val Phe Phe Gln Asn Ala Leu Ile
                    85                    90                    95
ttt ttt agc act ctg atc tac aaa ttt gga aga acc gaa gag cta tgg    542
Phe Phe Ser Thr Leu Ile Tyr Lys Phe Gly Arg Thr Glu Glu Leu Trp
                    100                    105                    110
acc tgagatcact tcttaagtca ctttttcctt ttgttatatt ctgtttgtag    595
Thr
atagggttttt tatctctcag tacacattgc caaatggagt agattgtaca ttaaattgtt    655
tgttttcttta cttttttatg ttctgagttt tgaaatagtt ttatgaaatt tctttatttt    715
tcattgcata gactgttaat atgtatataa tacaagacta tatgaattgg ataatgagta    775
tcagttttttt attcctgaga tttagaactt gatctactcc ctgagccagg gttacatcat    835
cttgtcattt tagaagtaac cactcttgct tctctggctg ggcacgggtg ctcattgcctg    895
taatcccagc actttgggag gccgaggcgg gccgattgct tgagggtcaag tgtttgagac    955
cagcctggcc aacatggcga aaccccatct actaaaaata caaaaattag ccaggcatgg    1015
tggtgggtgc ctgtaatccc aactacctag gaggctgagg caggagaatc gcttgaaccc    1075
gggggggcaga ggttgyagtg agctgagttt gcgccactgc actctagcct ggggggagaaa    1135
gtgaaactcc ctctcaaaaa aaaaaaamc    1164

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<210> 375

<211> 1250  
<212> DNA  
<213> Homo sapiens

<220>  
<221> CDS  
<222> 36..425

<221> sig\_peptide  
<222> 36..119  
<223> Von Heijne matrix  
score 11.6000003814697  
seq LLLLVQLLRFLRA/DG

<221> polyA\_signal  
<222> 1215..1220

<221> polyA\_site  
<222> 1240..1250

<400> 375

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atttcttccc cccgagctgg gcgtgcgcgg ccgca atg aac tgg gag ctg ctg      53
                                Met Asn Trp Glu Leu Leu
                                -25
ctg tgg ctg ctg gtg ctg tgc gcg ctg ctc ctg ctc ttg gtg cag ctg      101
Leu Trp Leu Leu Val Leu Cys Ala Leu Leu Leu Leu Leu Val Gln Leu
                                -20          -15          -10
ctg cgc ttc ctg agg gct gac ggc gac ctg acg cta cta tgg gcc gag      149
Leu Arg Phe Leu Arg Ala Asp Gly Asp Leu Thr Leu Leu Trp Ala Glu
                                -5          1          5          10
tgg cag gga cga cgc cca gaa tgg gag ctg act gat atg gtg gtg tgg      197
Trp Gln Gly Arg Arg Pro Glu Trp Glu Leu Thr Asp Met Val Val Trp
                                15          20          25
gtg act gga gcc tcg agt gga att ggt gag gag ctg gct tac cag ttg      245
Val Thr Gly Ala Ser Ser Gly Ile Gly Glu Glu Leu Ala Tyr Gln Leu
                                30          35          40
tct aaa cta gga gtt tct ctt gtg ctg tca gcc aga aga gtg cat gag      293
Ser Lys Leu Gly Val Ser Leu Val Leu Ser Ala Arg Arg Val His Glu
                                45          50          55
ctg gaa agg gtg aaa aga aga tgc cta gag aat ggc aat tta aaa gaa      341
Leu Glu Arg Val Lys Arg Arg Cys Leu Glu Asn Gly Asn Leu Lys Glu
                                60          65          70
aaa gat ata ctt gtt ttg ccc ctt gac ctg acc gac act ggt tcc cat      389
Lys Asp Ile Leu Val Leu Pro Leu Asp Leu Thr Asp Thr Gly Ser His
                                75          80          85          90
gaa agc ggc tac caa agc tgt tct cca gga att tgg tagaatcgac      435
Glu Ser Gly Tyr Gln Ser Cys Ser Pro Gly Ile Trp
                                95          100
attctggtca acaatgtgga aatgtcccag cggtctctgt gcatggatac caacttggat      495
gtctacagaa agctaagtga agcttaacta cttagggacg gtgtccttga caaatgtgk      555
kctgcctcac atgacgcaga ngaarcaagg aaagattgtt actgtgaata gcatcctggg      615
tatcatatct gtacctcttt ccattggata ctgtgctagc aagcatgctc tccggggkktk      675
ktttaatggc cttcraacag aacttgccac atacccargt ataatagttt ctaacatttg      735
cccaggacct gtgcaatcaa atattgtgga aaattcccta gctggagaag tcacaaagac      795
tataggcaat aatggagacc agtcccacaa gatgacaacc agtcgtttgt tgcggctgat      855
gttaatcagc atggccaatg atttgaaaga agtttgatc tcagaacaac ctttcttggt      915
agtaacatat ttgtggcaat acatgccaac ctgggcctgg tggataacca acaagatggg      975

```

```

gaagaaaagg attgagaact ttaagagtgg tgtggatgca gactcttctt attttaaaat 1035
ctttaagaca aaacatgact gaaaagagca cctgtacttt tcaagccact ggaggagaa 1095
atggaaaaca tgaaaacagc aatcttctta tgcttctgaa taatcaaaga ctaatttgtg 1155
attttacttt ttaatagata tgactttgct tccaacatgg aatgaaataa aaaataaata 1215
ataaaaagatt gccatgaatc ttgcaaaaaa aaaaa 1250

```

```

<210> 376
<211> 947
<212> DNA
<213> Homo sapiens

```

```

<220>
<221> CDS
<222> 155..751

```

```

<221> sig_peptide
<222> 155..340
<223> Von Heijne matrix
      score 3.70000004768372
      seq SILGIISVPLSIG/YC

```

```

<221> polyA_signal
<222> 912..917

```

```

<221> polyA_site
<222> 937..947

```

```

400> 376
agtgaaaaga agatgcctag agaatggcaa tttaaaagaa aaagatatac ttgttttgcc 60
ccttgacctg accgacactg gttcccatga agcggctacc aaagctgttc tccaggagtt 120
tggtagaatc gacattcttg tcaacaatgg tgga atg tcc cag cgt tct ctg tgc 175
                               Met Ser Gln Arg Ser Leu Cys
                               -60
atg gat acc agc ttg gat gtc tac aga rag cta ata gag ctt aac tac 223
Met Asp Thr Ser Leu Asp Val Tyr Arg Xaa Leu Ile Glu Leu Asn Tyr
55                               -50                               -45                               -40
tta ggg acg gtg tcc ttg aca aaa tgt gtt ctg cct cac atg atc gag 271
Leu Gly Thr Val Ser Leu Thr Lys Cys Val Leu Pro His Met Ile Glu
                               -35                               -30                               -25
agg aag caa gga aag att gtt act gtg aat agc atc ctg ggt atc ata 319
Arg Lys Gln Gly Lys Ile Val Thr Val Asn Ser Ile Leu Gly Ile Ile
                               -20                               -15                               -10
tct gta cct ctt tcc att gga tac tgt gct agc aag cat gct ctc cgg 367
Ser Val Pro Leu Ser Ile Gly Tyr Cys Ala Ser Lys His Ala Leu Arg
                               -5                               1                               5
ggt ttt ttt aat ggc ctt cga aca gaa ctt gcc aca tac cca ggt ata 415
Gly Phe Phe Asn Gly Leu Arg Thr Glu Leu Ala Thr Tyr Pro Gly Ile
10                               15                               20                               25
ata gtt tct aac att tgc cca gga cct gtg caa tca aat att gtg gaa 463
Ile Val Ser Asn Ile Cys Pro Gly Pro Val Gln Ser Asn Ile Val Glu
                               30                               35                               40
aat tcc cta gct gga gaa gtc aca aaa act ata ggc aat aat gga aac 511
Asn Ser Leu Ala Gly Glu Val Thr Lys Thr Ile Gly Asn Asn Gly Asn
                               45                               50                               55
cag tcc cac aag atg aca acc agt cgt tgt gtg cgg ctg atg tta atc 559

```

Gln	Ser	His	Lys	Met	Thr	Thr	Ser	Arg	Cys	Val	Arg	Leu	Met	Leu	Ile		
		60					65				70						
agc	atg	gcc	aat	gat	ttg	aaa	gaa	gtt	tg	atc	tca	gaa	caa	cct	ttc	607	
Ser	Met	Ala	Asn	Asp	Leu	Lys	Glu	Val	Trp	Ile	Ser	Glu	Gln	Pro	Phe		
		75				80					85						
ttg	tta	gta	aca	tat	ttg	tg	caa	tac	atg	cca	acc	tg	gcc	tg	tg	655	
Leu	Leu	Val	Thr	Tyr	Leu	Trp	Gln	Tyr	Met	Pro	Thr	Trp	Ala	Trp	Trp		
		90			95					100					105		
ata	acc	aac	aag	atg	ggg	aag	aaa	agg	att	gag	aac	ttt	aag	agt	gg	703	
Ile	Thr	Asn	Lys	Met	Gly	Lys	Lys	Arg	Ile	Glu	Asn	Phe	Lys	Ser	Gly		
			110					115				120					
gtg	gat	gcm	rac	tct	tct	tat	ttt	aaa	atc	ttt	aag	aca	aaa	cat	gac	751	
Val	Asp	Ala	Xaa	Ser	Ser	Tyr	Phe	Lys	Ile	Phe	Lys	Thr	Lys	His	Asp		
		125						130				135					
tgaaaaganc	acctgtactt	ttcaagccac	tgaggaggaga	aatggaaaac	atgaaaacag	811											
caatcttctt	atgcttctga	ataatcaaag	actaatttgt	gattttactt	tttaatagat	871											
atgactttgc	ttccaacatg	grrtgaaata	aaaaataaat	aataaaaagat	tgccatgrrt	931											
cttgcaaaaa	aaaaaa					947											

<210> 377  
 <211> 621  
 <212> DNA  
 <213> Homo sapiens  
 <220>  
 <221> CDS  
 <222> 46..585  
 <221> sig\_peptide  
 <222> 46..120  
 <223> Von Heijne matrix  
 score 6.30000019073486  
 seq AFSLSVMAALTFG/CF  
 <221> polyA\_signal  
 <222> 584..589  
 <221> polyA\_site  
 <222> 606..619

<400> 377																	
aactgggtgt	gcgtrtggag	tccggactcg	tgaggagacga	tcgcg	atg	aac	acg	gtg								57	
					Met	Asn	Thr	Val									
					-25												
ctg	tcg	cg	gc	aac	tca	ctg	ttc	gcc	ttc	tcg	ctg	agc	gtg	atg	gcs	105	
Leu	Ser	Arg	Ala	Asn	Ser	Leu	Phe	Ala	Phe	Ser	Leu	Ser	Val	Met	Ala		
	-20					-15				-10							
gcg	ctc	acc	ttc	ggc	tgc	ttc	atc	ayy	acc	gcc	ttc	aaa	gac	agg	agc	153	
Ala	Leu	Thr	Phe	Gly	Cys	Phe	Ile	Xaa	Thr	Ala	Phe	Lys	Asp	Arg	Ser		
-5				1				5					10				
gtc	ccg	gtg	cg	ctg	cac	gtc	tcg	cga	atc	atg	cta	aaa	aat	gta	gaa	201	
Val	Pro	Val	Arg	Leu	His	Val	Ser	Arg	Ile	Met	Leu	Lys	Asn	Val	Glu		
			15				20				25						
gat	ttc	act	gga	cct	aga	gaa	aga	agt	gat	ctg	gga	ttt	atc	aca	ttt	249	
Asp	Phe	Thr	Gly	Pro	Arg	Glu	Arg	Ser	Asp	Leu	Gly	Phe	Ile	Thr	Phe		

30	35	40	
gat ata act gct gat cta gag aat ata ttt gat tgg aat gtt aag cag			297
Asp Ile Thr Ala Asp Leu Glu Asn Ile Phe Asp Trp Asn Val Lys Gln			
45	50	55	
ttg ttt ctt tat tta tca gca gaa tat tca aca aaa aat aat gct ctg			345
Leu Phe Leu Tyr Leu Ser Ala Glu Tyr Ser Thr Lys Asn Asn Ala Leu			
60	65	70	75
aac caa ktt gtc cta tgg gac aag att gtt ttg aga ggt gat aat ccg			393
Asn Gln Xaa Val Leu Trp Asp Lys Ile Val Leu Arg Gly Asp Asn Pro			
80	85	90	
aag ctg ctg ctg aaa gat atg aaa aca aaa tat ttt ttc ttt gac gat			441
Lys Leu Leu Leu Lys Asp Met Lys Thr Lys Tyr Phe Phe Phe Asp Asp			
95	100	105	
gga aat ggt ctc wag gga aac agg aat gtc act ttg acc ctg tct tgg			489
Gly Asn Gly Leu Xaa Gly Asn Arg Asn Val Thr Leu Thr Leu Ser Trp			
110	115	120	
aac gtc gta cca aat gct gga att cta cct ctt gtg aca gga tca gga			537
Asn Val Val Pro Asn Ala Gly Ile Leu Pro Leu Val Thr Gly Ser Gly			
125	130	135	
cac gta tct gtc cca ttt cca gat aca tat gaa ata acg aag agt tat			585
His Val Ser Val Pro Phe Pro Asp Thr Tyr Glu Ile Thr Lys Ser Tyr			
140	145	150	155
caaatatttc tgaatttgaa acaaaaaaaaaaaaahm			621

<210> 378  
 <211> 52  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -20...-1

<400> 378  
 Met Pro Ser Val Asn Ser Ala Gly Leu Cys Val Leu Gln Leu Thr Thr  
 20 -15 -10 -5  
 Ala Val Thr Ser Ala Phe Leu Leu Ala Lys Val Asn Pro Phe Glu Xaa  
 1 5 10  
 Phe Leu Ser Arg Gly Phe Trp Leu Cys Ala Ala His His Phe Ile His  
 15 20 25  
 Pro Cys Leu Asp  
 30

<210> 379  
 <211> 193  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -23...-1

<400> 379

```

Met Val Val Leu Arg Ala Gly Lys Lys Thr Phe Leu Pro Pro Leu Xaa
      -20                      -15                      -10
Arg Ala Phe Ala Cys Arg Gly Cys Gln Leu Ala Pro Glu Arg Gly Ala
      -5                      1                      5
Glu Arg Arg Asp Thr Ala Pro Ser Gly Val Ser Arg Phe Cys Pro Pro
10      15      20      25
Arg Lys Ser Cys His Asp Trp Ile Gly Pro Pro Asp Lys Tyr Ser Asn
      30      35      40
Leu Arg Pro Val His Phe Tyr Ile Pro Glu Asn Glu Ser Pro Leu Glu
      45      50      55
Gln Lys Leu Arg Lys Leu Arg Gln Glu Thr Gln Glu Trp Asn Gln Gln
      60      65      70
Phe Trp Ala Asn Gln Asn Leu Thr Phe Ser Lys Glu Lys Glu Glu Phe
      75      80      85
Ile His Ser Arg Leu Lys Thr Lys Gly Leu Gly Leu Arg Thr Glu Ser
90      95      100      105
Gly Gln Lys Ala Thr Leu Asn Ala Glu Glu Met Ala Asp Phe Tyr Lys
      110      115      120
Glu Phe Leu Ser Lys Asn Phe Gln Lys His Met Tyr Tyr Asn Arg Asp
      125      130      135
Trp Tyr Lys Arg Asn Phe Ala Ile Thr Phe Phe Met Gly Lys Val Ala
      140      145      150
Leu Glu Arg Ile Trp Asn Lys Leu Lys Gln Lys Gln Lys Lys Arg Ser
155      160      165
Asn
170

```

```

<210> 380
<211> 82
<212> PRT
<213> Homo sapiens
<220>
<221> SIGNAL
<222> -14..-1

```

```

<400> 380
Met Ala Phe Thr Leu Xaa Ser Leu Leu Gln Ala Ala Leu Leu Cys Val
      -10                      -5                      1
Asn Ala Ile Ala Val Leu His Glu Glu Arg Phe Leu Lys Asn Ile Gly
      5      10      15
Trp Gly Thr Asp Gln Gly Ile Gly Gly Phe Gly Glu Glu Pro Gly Ile
      20      25      30
Lys Ser Xaa Xaa Met Xaa Leu Ile Arg Ser Val Arg Thr Val Met Arg
35      40      45      50
Val Pro Leu Ile Ile Val Asn Ser Ile Ala Ile Val Leu Leu Leu Leu
      55      60      65
Phe Gly

```

```

<210> 381
<211> 198
<212> PRT
<213> Homo sapiens

```

<220>

<221> SIGNAL

<222> -21..-1

<400> 381

```

Met Pro Val Pro Ala Leu Cys Leu Leu Trp Ala Leu Ala Met Val Thr
  -20                      -15                      -10
Arg Pro Ala Ser Ala Ala Pro Met Gly Gly Pro Glu Leu Ala Gln His
  -5                      1                      5                      10
Glu Glu Leu Thr Leu Leu Phe His Gly Thr Leu Gln Leu Gly Gln Ala
      15                      20                      25
Leu Asn Gly Val Tyr Arg Thr Thr Glu Gly Arg Leu Thr Lys Ala Arg
      30                      35                      40
Asn Ser Leu Gly Leu Tyr Gly Arg Thr Ile Glu Leu Leu Gly Gln Glu
      45                      50                      55
Val Ser Arg Gly Arg Asp Ala Ala Gln Glu Leu Arg Ala Ser Leu Leu
      60                      65                      70                      75
Glu Thr Gln Met Glu Glu Asp Ile Leu Xaa Leu Gln Ala Xaa Ala Thr
      80                      85                      90
Ala Glu Val Leu Gly Glu Val Ala Gln Ala Gln Lys Val Leu Arg Asp
      95                      100                      105
Ser Val Gln Arg Leu Xaa Xaa Gln Leu Xaa Xaa Ala Trp Leu Gly Pro
      110                      115                      120
Ala Tyr Arg Lys Phe Glu Val Leu Lys Ala Pro Pro Xaa Lys Gln Asn
      125                      130                      135
His Ile Leu Trp Ala Leu Thr Gly His Val Xaa Arg Gln Xaa Arg Glu
      140                      145                      150                      155
Met Val Ala Gln Gln Xaa Xaa Leu Xaa Gln Ile Gln Glu Lys Leu His
      160                      165                      170
Thr Ala Ala Leu Pro Ala
      175

```

<210> 382

<211> 160

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -55..-1

<400> 382

```

Met Asp Lys Leu Lys Lys Val Leu Ser Gly Gln Asp Thr Glu Asp Arg
  -55                      -50                      -45                      -40
Ser Gly Leu Ser Glu Val Val Glu Ala Ser Ser Leu Ser Trp Ser Thr
      -35                      -30                      -25
Arg Ile Lys Gly Phe Ile Ala Cys Phe Ala Ile Gly Ile Leu Cys Ser
      -20                      -15                      -10
Leu Leu Gly Thr Val Leu Leu Trp Val Pro Arg Lys Gly Leu His Leu
      -5                      1                      5
Phe Ala Val Phe Tyr Thr Phe Gly Asn Ile Ala Ser Ile Gly Ser Thr
      10                      15                      20                      25
Ile Phe Leu Met Gly Pro Val Lys Gln Leu Lys Arg Met Phe Glu Pro
      30                      35                      40

```



```

Thr Arg Leu Ile Ala Thr Ile Met Val Leu Leu Cys Phe Ala Leu Thr
      45                      50                      55
Leu Cys Ser Ala Phe Trp Trp His Asn Lys Gly Leu Ala Leu Ile Phe
      60                      65                      70
Cys Ile Leu Gln Ser Leu Ala Leu Thr Trp Tyr Ser Leu Ser Phe Ile
      75                      80                      85
Pro Phe Ala Arg Asp Ala Val Lys Xaa Cys Phe Ala Val Cys Leu Ala
      90                      95                      100                      105

```

<210> 383  
 <211> 108  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -18...-1

<400> 383

```

Met Lys Ala Leu Cys Leu Leu Leu Leu Pro Val Leu Gly Leu Leu Val
      -15                      -10                      -5
Ser Ser Lys Thr Leu Cys Ser Met Glu Glu Ala Ile Asn Glu Arg Ile
      1                      5                      10
Gln Glu Val Ala Gly Ser Leu Ile Phe Arg Ala Ile Ser Ser Ile Gly
      15                      20                      25                      30
Arg Gly Ser Glu Ser Val Thr Ser Arg Gly Asp Leu Ala Thr Cys Pro
      35                      40                      45
Arg Gly Phe Ala Val Thr Gly Cys Thr Cys Gly Ser Ala Cys Gly Ser
      50                      55                      60
Trp Asp Val Arg Ala Glu Thr Thr Cys His Cys Gln Cys Ala Gly Met
      65                      70                      75
Asp Trp Thr Gly Ala Arg Cys Cys Arg Val Gln Pro
      80                      85                      90

```

<210> 384  
 <211> 64  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -22...-1

<400> 384

```

Met Ile Ser Arg Gln Leu Arg Ser Leu Ser Cys Leu Cys Pro Ala Leu
      -20                      -15                      -10
Phe Pro Gly Thr Ser Ser Phe Ile Val Ala Leu Ser Ser Pro Ala Asp
      -5                      1                      5                      10
Leu Tyr Ile Pro Xaa Arg Xaa Arg Ser Asp Glu Leu Val Phe Glu Ser
      15                      20                      25
Gln Lys Gly Ser Ala Met Glu Leu Ala Val Ile Thr Val Xaa Gly Val
      30                      35                      40

```

<210> 385  
 <211> 27  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -15..-1

<400> 385  
 Met Gly Phe Leu Xaa Leu Met Thr Leu Thr Thr His Val His Ser Ser  
 -15 -10 -5 1  
 Ala Lys Pro Asn Glu Gln Pro Trp Leu Leu Asn  
 5 10

<210> 386  
 <211> 186  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -21..-1

<400> 386  
 Met Ser Pro Ser Gly Arg Leu Cys Leu Leu Thr Ile Val Gly Leu Ile  
 -20 -15 -10  
 Leu Pro Thr Arg Gly Gln Thr Leu Lys Asp Thr Thr Ser Ser Ser Ser  
 5 1 5 10  
 Ala Asp Ser Thr Ile Met Asp Ile Gln Val Pro Thr Arg Ala Pro Asp  
 15 20 25  
 Ala Val Tyr Thr Glu Leu Gln Pro Thr Ser Pro Thr Pro Thr Trp Pro  
 30 35 40  
 Ala Asp Glu Thr Pro Gln Pro Gln Thr Gln Thr Gln Gln Leu Glu Gly  
 45 50 55  
 Thr Asp Gly Pro Leu Val Thr Asp Pro Glu Thr His Xaa Ser Xaa Lys  
 60 65 70 75  
 Ala Ala His Pro Thr Asp Asp Thr Thr Thr Leu Ser Glu Arg Pro Ser  
 80 85 90  
 Pro Ser Thr Xaa Val His Xaa Arg Pro Xaa Xaa Pro Ser Xaa His Leu  
 95 100 105  
 Val Phe Met Arg Met Thr Pro Ser Ser Met Met Asn Thr Pro Ser Gly  
 110 115 120  
 Asn Xaa Gly Cys Trp Ser Gln Leu Cys Cys Ser Ser Gln Ala Ser Ser  
 125 130 135  
 Ser Ser Pro Val Ala Ser Ala Gly Ser Cys Pro Gly Tyr Ala Gly Ile  
 140 145 150 155  
 Ile Ala Gly Glu Ser Ile Arg Asn Arg Ser  
 160 165

<210> 387

<211> 179  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -26..-1

<400> 387

```
Met Glu Thr Gly Ala Leu Arg Arg Pro Gln Leu Leu Pro Leu Leu Leu
  -25                -20                -15
Leu Leu Cys Gly Pro Ser Gln Asp Gln Cys Arg Pro Val Leu Gln Asn
  -10                -5                1                5
Leu Leu Gln Ser Pro Gly Leu Thr Trp Ser Leu Glu Val Pro Thr Gly
          10                15                20
Arg Glu Gly Lys Glu Gly Gly Asp Arg Gly Pro Gly Leu Xaa Gly Ala
      25                30                35
Thr Pro Ala Arg Ser Pro Gln Gly Lys Glu Met Gly Arg Gln Arg Thr
      40                45                50
Arg Lys Val Lys Gly Pro Ala Trp Xaa His Thr Ala Asn Gln Glu Leu
  55                60                65                70
Asn Arg Met Arg Ser Leu Ser Ser Gly Ser Val Pro Val Gly His Leu
          75                80                85
Glu Gly Gly Thr Val Lys Leu Gln Lys Asp Thr Gly Leu His Ser Cys
          90                95                100
Xaa Asp Gly Met Ala Ser Leu Glu Gly Thr Pro Ala Ser Val Leu Ala
      105                110                115
Asp Ala Cys Pro Gly Phe His Asp Val Xaa Val Gln Xaa Ala Leu Phe
      120                125                130
Gly Leu Ser Gly Xaa Xaa Leu Trp Leu Lys Thr His Phe Cys Leu Ser
  135                140                145                150
Ile Xaa Leu
```

<210> 388  
<211> 150  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -55..-1

<400> 388

```
Met Ala Thr Thr Val Pro Asp Gly Cys Arg Asn Gly Leu Lys Ser Lys
  -55                -50                -45                -40
Tyr Tyr Arg Leu Cys Asp Lys Ala Glu Ala Trp Gly Ile Val Leu Glu
          -35                -30                -25
Thr Val Ala Thr Ala Gly Val Val Thr Ser Val Ala Phe Met Leu Thr
          -20                -15                -10
Leu Pro Ile Leu Val Cys Lys Val Gln Asp Ser Asn Arg Arg Lys Met
      -5                1                5
Leu Pro Thr Gln Phe Leu Phe Leu Leu Gly Val Leu Gly Ile Phe Gly
  10                15                20                25
Leu Thr Phe Ala Phe Ile Ile Gly Leu Asp Gly Ser Thr Gly Pro Thr
          30                35                40
```

```

Arg Phe Phe Leu Phe Gly Ile Leu Phe Ser Ile Cys Phe Ser Cys Leu
      45                      50                      55
Leu Ala His Ala Val Ser Leu Thr Lys Leu Val Arg Gly Arg Lys Ala
      60                      65                      70
Pro Phe Pro Val Gly Asp Ser Gly Ser Gly Arg Gly Leu Gln Pro Ser
      75                      80                      85
Pro Gly Cys Tyr Arg Tyr
90                      95

```

<210> 389  
 <211> 236  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -31...-1

<400> 389

```

Met Leu Ser Lys Gly Leu Lys Arg Lys Arg Glu Glu Glu Glu Lys
-30                      -25                      -20
Glu Pro Leu Ala Val Asp Ser Trp Trp Leu Asp Pro Gly His Ala Ala
15                      -10                      -5                      1
Val Ala Gln Ala Pro Pro Ala Val Ala Ser Ser Ser Leu Phe Asp Leu
      5                      10                      15
Ser Val Leu Lys Leu His His Ser Leu Gln Xaa Ser Xaa Pro Asp Leu
      20                      25                      30
Arg His Leu Val Leu Val Xaa Asn Thr Leu Arg Arg Ile Gln Ala Ser
      35                      40                      45
Met Ala Pro Ala Ala Ala Leu Pro Pro Val Pro Thr Pro Pro Ala Ala
50                      55                      60                      65
Pro Xaa Val Ala Asp Asn Leu Leu Ala Ser Ser Asp Ala Ala Leu Ser
      70                      75                      80
Ala Ser Met Ala Xaa Leu Leu Glu Asp Leu Ser His Ile Glu Gly Leu
      85                      90                      95
Ser Gln Ala Pro Gln Pro Leu Ala Asp Glu Gly Pro Pro Gly Arg Ser
100                      105                      110
Ile Gly Gly Xaa Pro Pro Xaa Leu Gly Ala Leu Asp Leu Leu Gly Pro
115                      120                      125
Ala Thr Gly Cys Leu Leu Asp Asn Gly Leu Glu Gly Leu Phe Glu Asp
130                      135                      140                      145
Ile Asp Thr Ser Met Tyr Asp Asn Glu Leu Trp Ala Pro Ala Ser Glu
      150                      155                      160
Gly Leu Lys Pro Gly Pro Glu Asp Gly Pro Gly Lys Glu Glu Ala Pro
165                      170                      175
Glu Leu Asp Glu Ala Glu Leu Asp Tyr Leu Met Asp Val Leu Val Gly
180                      185                      190
Thr Gln Ala Leu Glu Arg Pro Pro Gly Pro Gly Arg
195                      200                      205

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<210> 390  
 <211> 149  
 <212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -100..-1

<400> 390

```

Met Glu Thr Leu Tyr Arg Val Pro Phe Leu Val Leu Glu Cys Pro Asn
-100                               -95             -90             -85
Leu Lys Leu Lys Lys Pro Pro Trp Leu His Met Pro Ser Ala Met Thr
                               -80             -75             -70
Val Tyr Ala Leu Val Val Val Ser Tyr Phe Leu Ile Thr Gly Gly Ile
                               -65             -60             -55
Ile Tyr Asp Val Ile Val Glu Pro Pro Ser Val Gly Ser Met Thr Asp
                               -50             -45             -40
Glu His Gly His Gln Arg Pro Val Ala Phe Leu Ala Tyr Arg Val Asn
                               -35             -30             -25
Gly Gln Tyr Ile Met Glu Gly Leu Ala Ser Ser Phe Leu Phe Thr Met
-20                               -15             -10             -5
Gly Gly Leu Gly Phe Ile Ile Leu Asp Gly Ser Asn Ala Pro Asn Ile
                               1             5             10
Pro Lys Leu Asn Arg Phe Leu Leu Leu Phe Ile Gly Phe Val Cys Val
                               15             20             25
Leu Xaa Ser Phe Xaa Xaa Ala Arg Val Phe Met Arg Met Lys Leu Pro
                               30             35             40
Gly Tyr Leu Met Gly
45

```

<210> 391

<211> 69

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -49..-1

<400> 391

```

Met Pro Phe His Phe Pro Phe Leu Gly Phe Val Cys Leu His Leu His
                               -45             -40             -35
Leu Thr Pro Cys Leu Thr Val Pro Arg Arg Pro Leu Phe Leu Leu Leu
                               -30             -25             -20
His Leu Cys Pro His Leu Pro Phe Leu Leu Leu Leu Ser Cys Val Gly
                               -15             -10             -5
Xaa Xaa Pro Ser Cys Leu Pro Ser Ser Ser Thr Cys Val Ser Leu His
                               1             5             10             15
Phe Phe Ile Pro Asp
                               20

```

<210> 392

<211> 241

<212> PRT

<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -30..-1

<400> 392

Met Gly Thr Ala Ser Arg Ser Asn Ile Ala Arg His Leu Gln Thr Asn  
-30 -25 -20 -15  
Leu Ile Leu Phe Cys Val Gly Ala Val Gly Ala Cys Thr Leu Ser Val  
-10 -5 1  
Thr Gln Pro Trp Tyr Leu Glu Val Asp Tyr Thr His Glu Ala Val Thr  
5 10 15  
Ile Lys Cys Thr Phe Ser Ala Thr Gly Cys Pro Ser Glu Gln Pro Thr  
20 25 30  
Cys Leu Trp Phe Arg Tyr Gly Ala His Gln Pro Glu Asn Leu Cys Leu  
35 40 45 50  
Asp Gly Cys Lys Ser Glu Ala Xaa Lys Phe Thr Val Arg Glu Ala Leu  
55 60 65  
Lys Glu Asn Gln Val Ser Leu Thr Val Asn Arg Val Thr Ser Asn Asp  
70 75 80  
Ser Ala Ile Tyr Ile Cys Gly Ile Ala Phe Pro Ser Val Pro Glu Ala  
85 90 95  
Arg Ala Lys Gln Thr Gly Gly Gly Thr Thr Leu Val Val Arg Glu Ile  
100 105 110  
Lys Leu Leu Ser Lys Glu Leu Arg Ser Phe Leu Thr Ala Leu Val Ser  
115 120 125 130  
Leu Leu Ser Val Tyr Val Thr Gly Val Cys Val Ala Phe Ile Leu Leu  
135 140 145  
Ser Lys Ser Lys Ser Asn Pro Leu Arg Asn Lys Glu Ile Lys Glu Asp  
150 155 160  
Ser Gln Lys Lys Lys Ser Ala Arg Arg Ile Phe Gln Glu Ile Ala Gln  
165 170 175  
Glu Leu Tyr His Lys Arg His Val Glu Thr Asn Gln Gln Ser Glu Lys  
180 185 190  
Asp Asn Asn Thr Tyr Glu Asn Arg Arg Val Leu Ser Asn Tyr Glu Arg  
195 200 205 210  
Pro

<210> 393  
<211> 47  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -30..-1

<400> 393

Met Asn Cys Asn Val Val Ser Glu Arg Gly Lys Trp Leu Glu Val Glu  
-30 -25 -20 -15  
Cys Ser Leu Met Thr Cys Thr Thr Leu Ile Asn Ala Ser Ala Ile Ser  
-10 -5 1  
Thr Asn Thr Leu Thr Asp Met Gly Ser Phe Asp Arg Arg Glu Ser  
5 10 15

<210> 394  
 <211> 65  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -28..-1

<400> 394  
 Met Ala Phe Gly Leu Gln Met Phe Ile Gln Arg Lys Phe Pro Tyr Pro  
                   -25                  -20                  -15  
 Leu Gln Trp Ser Leu Leu Val Ala Val Val Ala Gly Ser Val Val Ser  
           -10                  -5                  1  
 Tyr Gly Val Thr Arg Val Glu Ser Glu Lys Cys Asn Asn Leu Trp Leu  
 5                  10                  15                  20  
 Phe Leu Glu Thr Gly Gln Leu Pro Lys Asp Arg Ser Thr Asp Gln Xaa  
                   25                  30                  35  
 Ser

<210> 395  
 <211> 73  
 <212> PRT  
 <213> Homo sapiens  
 <220>  
 <221> SIGNAL  
 <222> -24..-1

<400> 395  
 Met Thr Cys Trp Met Leu Pro Pro Ile Ser Phe Leu Ser Tyr Leu Pro  
                   -20                  -15                  -10  
 Leu Trp Leu Gly Pro Ile Trp Pro Cys Ser Gly Ser Thr Leu Gly Lys  
           -5                  1                  5  
 Pro Asp Pro Gly Val Trp Pro Ser Leu Phe Arg Pro Trp Asp Ala Ala  
 10                  15                  20  
 Ser Pro Gly Asn Tyr Ala Leu Ser Arg Gly Xaa Asn Xaa Tyr Xaa Xaa  
 25                  30                  35                  40  
 Trp Gly Gln Gly Thr His Ser Ser Leu  
                   45

<210> 396  
 <211> 60  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -18..-1

<400> 396

Met	Pro	Cys	Pro	Thr	Trp	Thr	Cys	Leu	Lys	Ser	Phe	Pro	Ser	Pro	Thr
			-15					-10					-5		
Ser	Ser	His	Ala	Ser	Ser	Leu	His	Leu	Pro	Pro	Ser	Cys	Thr	Arg	Leu
		1				5					10				
Thr	Leu	Thr	Gln	Thr	Leu	Arg	Thr	Gly	Met	His	Leu	Ser	Arg	Ala	Leu
15					20					25					30
Gln	Gly	Thr	Leu	Thr	Arg	Leu	Gln	Ser	Thr	Pro	Ala				
				35					40						

<210> 397  
 <211> 192  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -93...-1

<400> 397

Met	Ala	Glu	Leu	Gly	Leu	Asn	Glu	His	His	Gln	Asn	Glu	Val	Ile	Asn
			-90					-85					-80		
Tyr	Met	Arg	Phe	Ala	Arg	Ser	Lys	Arg	Gly	Leu	Arg	Leu	Lys	Thr	Val
		-75					-70					-65			
Asp	Ser	Cys	Phe	Gln	Asp	Leu	Lys	Glu	Ser	Arg	Leu	Val	Glu	Asp	Thr
		-60				-55					-50				
Phe	Thr	Ile	Asp	Glu	Val	Ser	Glu	Val	Leu	Asn	Gly	Leu	Gln	Ala	Val
45				-40						-35					-30
Val	His	Ser	Glu	Val	Glu	Ser	Glu	Leu	Ile	Asn	Thr	Ala	Tyr	Thr	Asn
			-25					-20						-15	
Val	Leu	Leu	Leu	Arg	Gln	Leu	Phe	Ala	Gln	Ala	Glu	Lys	Trp	Tyr	Leu
		-10					-5					1			
Lys	Leu	Gln	Thr	Asp	Ile	Ser	Glu	Leu	Glu	Asn	Arg	Glu	Leu	Leu	Glu
	5				10					15					
Gln	Xaa	Ala	Glu	Phe	Glu	Lys	Ala	Xaa	Ile	Thr	Ser	Ser	Asn	Lys	Lys
20					25					30					35
Pro	Ile	Leu	Xaa	Val	Thr	Xaa	Pro	Lys	Leu	Ala	Pro	Leu	Asn	Glu	Gly
			40					45						50	
Gly	Thr	Ala	Lys	Leu	Leu	Asn	Lys	Val	Ile	Cys	Ile	Ile	Leu	Arg	Asn
		55					60					65			
Gly	Lys	Ser	Leu	Ile	Leu	Ser	Cys	His	Cys	Leu	Gly	Trp	Arg	Asn	Lys
	70					75					80				
Ser	Gly	Arg	Phe	Val	Ser	Gly	Pro	Leu	Arg	Ile	Ile	Ser	Pro	Leu	Gln
85						90					95				

<210> 398  
 <211> 149  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -72...-1



<400> 398

```

Met Asn Leu Phe Ile Met Tyr Met Ala Gly Asn Thr Ile Ser Ile Phe
   -70                      -65                      -60
Pro Thr Met Met Val Cys Met Met Ala Trp Arg Pro Ile Gln Ala Leu
   -55                      -50                      -45
Met Ala Ile Ser Ala Thr Phe Lys Met Leu Glu Ser Ser Ser Gln Lys
   -40                      -35                      -30                      -25
Phe Leu Gln Gly Leu Val Tyr Leu Ile Gly Asn Leu Met Gly Leu Ala
                      -20                      -15                      -10
Leu Ala Val Tyr Lys Cys Gln Ser Met Gly Leu Leu Pro Thr His Ala
                      -5                      1                      5
Ser Asp Trp Leu Ala Phe Ile Glu Pro Pro Glu Arg Met Glu Ser Val
   10                      15                      20
Val Glu Asp Cys Phe Cys Glu His Glu Lys Ala Ala Pro Gly Pro Tyr
   25                      30                      35                      40
Val Phe Gly Ser Tyr Leu His Pro Ser Leu Ser Pro Val Ala Pro Gln
                      45                      50                      55
His Thr Leu Lys Leu Ile Thr Tyr Val Lys Lys Asn Gln Lys Thr Leu
                      60                      65                      70
Phe Ser Met Val Gly
                      75

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<210> 399

<211> 73

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -20..-1

<400> 399

```

Met Thr Pro Leu Leu Thr Leu Ile Leu Val Val Leu Met Gly Leu Pro
   20                      -15                      -10                      -5
Leu Ala Gln Ala Leu Asp Cys His Val Cys Ala Tyr Asn Gly Asp Asn
                      1                      5                      10
Cys Phe Asn Pro Met Arg Cys Pro Ala Met Val Ala Tyr Cys Met Thr
   15                      20                      25
Thr Arg Thr Tyr Tyr Thr Pro Thr Arg Met Lys Val Ser Lys Ser Cys
   30                      35                      40
Val Pro Arg Cys Phe Glu Xaa Cys Val
   45                      50

```

<210> 400

<211> 86

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -20..-1

<400> 400

```

Met Asn Leu His Phe Pro Gln Trp Phe Val His Ser Ser Ala Leu Gly
-20          -15          -10          -5
Leu Val Leu Ala Pro Pro Phe Ser Ser Pro Gly Thr Asp Pro Thr Phe
          1          5          10
Pro Cys Ile Tyr Cys Arg Leu Leu Asn Met Ile Met Thr Arg Leu Ala
          15          20          25
Phe Ser Phe Ile Thr Cys Leu Cys Pro Asn Leu Lys Glu Val Cys Leu
          30          35          40
Ile Leu Pro Glu Lys Asn Cys Asn Ser Arg His Ala Gly Phe Val Gly
45          50          55          60
Pro Xaa Lys Leu Arg Gln
          65

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<210> 401  
 <211> 78  
 <212> PRT  
 <213> Homo sapiens

<220>

<221> SIGNAL

<222> -21..-1

<400> 401

```

Met Cys Pro Val Phe Ser Lys Gln Leu Leu Ala Cys Gly Ser Leu Leu
-20          -15          -10
Pro Gly Leu Trp Gln His Leu Thr Ala Asn His Trp Pro Pro Phe Ser
5          1          5          10
Xaa Phe Leu Cys Thr Val Cys Ser Gly Ser Ser Glu Gln Ile Ser Glu
          15          20          25
Tyr Thr Ala Ser Ala Thr Pro Pro Leu Cys Arg Ser Leu Asn Gln Glu
          30          35          40
Pro Phe Val Ser Arg Ala Ile Arg Pro Lys Tyr Ser Ile Thr
45          50          55

```

<210> 402  
 <211> 65  
 <212> PRT  
 <213> Homo sapiens

<220>

<221> SIGNAL

<222> -28..-1

<400> 402

```

Met Gly Lys Gly His Gln Arg Pro Trp Trp Lys Val Leu Pro Leu Ser
          -25          -20          -15
Cys Phe Leu Val Ala Leu Ile Ile Trp Cys Tyr Leu Arg Glu Glu Ser
          -10          -5          1
Glu Ala Asp Gln Trp Leu Arg Gln Val Trp Gly Glu Val Pro Glu Pro
5          10          15          20
Ser Asp Arg Ser Glu Glu Pro Glu Thr Pro Ala Ala Tyr Arg Ala Arg
          25          30          35
Thr

```

<210> 403  
 <211> 211  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -27...-1

<400> 403

Met	Leu	Leu	Leu	Ser	Ile	Thr	Thr	Ala	Tyr	Thr	Gly	Leu	Glu	Leu	Thr
	-25						-20					-15			
Phe	Phe	Ser	Gly	Val	Tyr	Gly	Thr	Cys	Ile	Gly	Ala	Thr	Asn	Lys	Phe
	-10				-5					1				5	
Gly	Ala	Glu	Glu	Xaa	Ser	Leu	Ile	Gly	Leu	Ser	Gly	Ile	Phe	Ile	Gly
			10					15					20		
Ile	Gly	Glu	Ile	Leu	Gly	Gly	Ser	Leu	Phe	Gly	Leu	Leu	Ser	Lys	Asn
			25				30						35		
Asn	Arg	Phe	Gly	Arg	Asn	Pro	Val	Val	Leu	Leu	Gly	Ile	Leu	Val	His
	40					45					50				
Phe	Ile	Ala	Phe	Tyr	Leu	Ile	Phe	Leu	Asn	Met	Pro	Gly	Asp	Ala	Pro
	55				60					65					
Ile	Ala	Pro	Val	Lys	Gly	Thr	Asp	Ser	Ser	Ala	Tyr	Ile	Lys	Ser	Ser
	70			75						80				85	
Lys	Xaa	Phe	Ala	Ile	Leu	Cys	Xaa	Phe	Leu	Xaa	Gly	Leu	Gly	Asn	Ser
			90				95						100		
Cys	Phe	Asn	Thr	Xaa	Leu	Leu	Xaa	Ile	Xaa	Gly	Phe	Leu	Tyr	Ser	Glu
		105					110					115			
Xaa	Ser	Ala	Pro	Xaa	Phe	Ala	Ile	Phe	Asn	Phe	Val	Gln	Ser	Ile	Cys
	120					125						130			
Ala	Ala	Val	Ala	Phe	Phe	Tyr	Ser	Asn	Tyr	Leu	Leu	Leu	His	Trp	Gln
	135				140						145				
Leu	Leu	Val	Met	Val	Ile	Phe	Gly	Phe	Xaa	Gly	Thr	Ile	Ser	Phe	Phe
	150			155					160					165	
Thr	Val	Glu	Trp	Glu	Xaa	Ala	Ala	Phe	Val	Xaa	Arg	Gly	Ser	Asp	Tyr
			170					175						180	

Arg Ser Ile

<210> 404  
 <211> 123  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -80...-1

<400> 404

Met	Ser	Thr	Trp	Tyr	Leu	Ala	Leu	Asn	Lys	Ser	Tyr	Lys	Asn	Lys	Asp
-80				-75				-70					-65		
Ser	Val	Arg	Ile	Tyr	Leu	Ser	Leu	Cys	Thr	Val	Ser	Ile	Lys	Phe	Thr
			-60					-55					-50		

Tyr Phe His Asp Ile Gln Thr Asn Cys Leu Thr Thr Trp Lys His Ser  
 -45 -40 -35  
 Arg Cys Arg Phe Tyr Trp Ala Phe Gly Gly Ser Ile Leu Gln His Ser  
 -30 -25 -20  
 Val Asp Pro Leu Val Leu Phe Leu Ser Leu Ala Leu Leu Val Thr Pro  
 -15 -10 -5  
 Thr Ser Thr Pro Ser Ala Lys Ile Gln Ser Leu Gln Ile Asp Leu Pro  
 1 5 10 15  
 Gly Gly Trp Arg Leu Ala Thr Asp Arg Ile Phe Thr Leu Ser Pro Val  
 20 25 30  
 Pro Met Asp Xaa Pro Leu Ile Leu His Gln Leu  
 35 40

<210> 405  
 <211> 86  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -26...-1

<400> 405  
 Met Glu Lys Ser Trp Met Leu Trp Asn Phe Val Glu Arg Trp Leu Ile  
 -25 -20 -15  
 Ala Leu Ala Ser Trp Ser Trp Ala Leu Cys Arg Ile Ser Leu Leu Pro  
 10 -5 1 5  
 Leu Ile Val Thr Phe His Leu Tyr Gly Gly Ile Ile Leu Leu Leu Leu  
 10 15 20  
 Ile Phe Ile Ser Ile Xaa Gly Ile Leu Tyr Lys Phe Xaa Asp Val Leu  
 25 30 35  
 Leu Tyr Phe Pro Xaa Gln Xaa Ser Ser Ser Arg Leu Tyr Asp Ser His  
 40 45 50  
 Ala His Trp Xaa Ser Xaa  
 55 60

<210> 406  
 <211> 162  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -31...-1

<400> 406  
 Met Ala Ala Ala Trp Pro Ser Gly Pro Xaa Ala Pro Glu Ala Val Thr  
 -30 -25 -20  
 Ala Arg Leu Val Gly Val Leu Trp Phe Val Ser Val Thr Thr Gly Pro  
 -15 -10 -5 1  
 Trp Gly Ala Val Ala Thr Ser Ala Gly Gly Glu Glu Ser Leu Lys Cys  
 5 10 15  
 Glu Asp Leu Lys Val Gly Gln Tyr Ile Cys Lys Asp Pro Lys Ile Asn

20 25 30  
Asp Ala Thr Gln Glu Pro Val Asn Cys Thr Asn Tyr Thr Ala His Val  
35 40 45  
Ser Cys Phe Pro Ala Pro Asn Ile Thr Cys Lys Asp Ser Ser Gly Asn  
50 55 60 65  
Glu Thr His Phe Thr Gly Asn Glu Val Gly Phe Phe Lys Pro Ile Ser  
70 75 80  
Cys Arg Asn Val Asn Gly Tyr Ser Tyr Asn Glu Gln Ser His Val Ser  
85 90 95  
Phe Ser Trp Met Val Gly Ser Arg Ser Ile Leu Pro Trp Ile Pro Cys  
100 105 110  
Phe Gly Phe Val Lys Xaa Xaa His Cys Arg Val Xaa Trp Asn Trp Glu  
115 120 125  
Pro Asn  
130

<210> 407  
<211> 98  
<212> PRT  
<213> Homo sapiens  
<220>  
<221> SIGNAL  
<222> -37...-1

<400> 407  
Met Ala Ser Leu Leu Cys Cys Gly Pro Lys Leu Ala Ala Cys Gly Ile  
-35 -30 -25  
Val Leu Ser Ala Trp Gly Val Ile Met Leu Ile Met Leu Gly Ile Phe  
-20 -15 -10  
Phe Asn Val His Ser Ala Val Leu Ile Glu Asp Val Pro Phe Thr Glu  
5 1 5 10  
Lys Asp Phe Glu Asn Gly Pro Gln Asn Ile Tyr Asn Leu Tyr Xaa Gln  
15 20 25  
Xaa Ser Tyr Asn Cys Phe Ile Ala Ala Gly Leu Tyr Leu Leu Gly  
30 35 40  
Gly Phe Ser Phe Cys Gln Xaa Arg Leu Asn Lys Arg Lys Glu Tyr Met  
45 50 55  
Val Arg  
60

<210> 408  
<211> 70  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -15...-1

<400> 408  
Met Arg Phe Leu Pro Cys Cys Leu Leu Trp Ser Val Phe Asn Pro Glu  
-15 -10 -5 1

Ser Leu Asn Cys His Tyr Phe Xaa Xaa Glu Xaa Cys Ile Phe Xaa Ser  
5 10 15  
Leu Gln Tyr Tyr Glu Ile Ser Leu Gln Glu Lys Leu Leu Gly Phe Leu  
20 25 30  
Trp Leu Cys Phe Leu Ser Tyr Phe Phe Arg Ala Val Tyr Phe Leu Ile  
35 40 45  
Asp Phe Ser Ser Phe Thr  
50 55

<210> 409  
<211> 60  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -45..-1

<400> 409

Met His Ser Leu Phe Ile Ala Ser Leu Lys Val Leu Phe Tyr Tyr Ser  
45 -40 -35 -30  
Phe Ser Phe Arg Phe Asn Trp Phe Asp Cys Leu Leu His Asn Leu Gly  
-25 -20 -15  
Glu Asn Phe Leu Ser Leu Leu Ser Lys Ser Cys Ser Ala Asp Pro Ser  
-10 -5 1  
Gly Ser Thr Phe Met Arg Asp Ile Glu Thr Asn Lys  
5 10 15

<210> 410  
<211> 39  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -22..-1

<400> 410

Met Pro Glu Ala Val Glu Gln Ser Ala His Leu Phe Val Thr Trp Ser  
-20 -15 -10  
Ser Gln Arg Ala Leu Ser His Pro Ala Pro Phe Leu Thr Xaa Xaa Lys  
-5 1 5 10  
Asn Pro Phe Leu Trp Lys Leu  
15

<210> 411  
<211> 51  
<212> PRT  
<213> Homo sapiens

<220>

<221> SIGNAL  
<222> -23..-1

<400> 411  
Met Ala Phe Gln Ser Leu Leu Glu Met Lys Phe Phe Leu Cys Ala Ala  
          -20                  -15                  -10  
Phe Pro Leu Gly Ala Gly Val Lys Met Phe His Tyr Leu Gly Pro Gly  
          -5                  1                  5  
Lys Pro Leu Xaa Gln Ala Ser Pro Ser Pro His Pro His Arg Xaa Arg  
10                  15                  20                  25  
Ile Trp Pro

<210> 412  
<211> 95  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -48..-1

<400> 412  
Met Ala Ser Ser His Trp Asn Glu Thr Thr Thr Ser Val Tyr Gln Tyr  
          -45                  -40                  -35  
Leu Gly Phe Gln Val Gln Lys Ile Tyr Pro Phe His Asp Asn Trp Asn  
          -30                  -25                  -20  
Thr Ala Cys Phe Val Ile Leu Leu Leu Phe Ile Phe Thr Val Val Ser  
          -15                  -10                  -5  
Leu Val Val Leu Ala Phe Leu Tyr Glu Val Leu Xaa Xaa Cys Cys Cys  
1                  5                  10                  15  
Val Lys Asn Lys Thr Val Lys Asp Leu Lys Ser Glu Pro Asn Pro Leu  
          20                  25                  30  
Xaa Xaa Met Met Asp Asn Ile Arg Lys Arg Glu Thr Glu Val Val  
          35                  40                  45

<210> 413  
<211> 60  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -32..-1

<400> 413  
Met Asp Glu Tyr Ser Trp Trp Cys His Val Leu Glu Val Val Lys Gly  
          -30                  -25                  -20  
Gln Met Phe Thr Phe Ile Asn Ile Thr Leu Trp Leu Gly Ser Leu Cys  
          -15                  -10                  -5  
Gln Arg Phe Phe Tyr Ala Ser Gly Thr Tyr Phe Leu Ile Tyr Ile Ser  
1                  5                  10                  15  
Thr Val Thr Pro Ser Trp Arg Leu Cys Leu Val Ser  
          20                  25

<210> 414  
 <211> 170  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -79...-1

<400> 414

Met	Glu	Asp	Pro	Asn	Pro	Glu	Glu	Asn	Met	Lys	Gln	Gln	Asp	Ser	Pro
				-75					-70					-65	
Lys	Glu	Arg	Ser	Pro	Gln	Ser	Pro	Gly	Gly	Asn	Ile	Cys	His	Leu	Gly
			-60					-55					-50		
Ala	Pro	Lys	Cys	Thr	Arg	Cys	Leu	Ile	Thr	Phe	Ala	Asp	Ser	Lys	Phe
		-45					-40					-35			
Gln	Glu	Arg	His	Met	Lys	Arg	Glu	His	Pro	Ala	Asp	Phe	Val	Ala	Gln
	-30				-25						-20				
Lys	Leu	Gln	Gly	Val	Leu	Phe	Ile	Cys	Phe	Thr	Cys	Ala	Arg	Ser	Phe
15				-10					-5						1
Pro	Ser	Ser	Lys	Ala	Xaa	Xaa	Thr	His	Gln	Arg	Ser	His	Gly	Pro	Xaa
		5					10					15			
Ala	Lys	Pro	Thr	Leu	Pro	Val	Ala	Thr	Thr	Thr	Ala	Gln	Pro	Thr	Phe
		20					25					30			
Pro	Cys	Pro	Asp	Cys	Gly	Lys	Thr	Phe	Gly	Gln	Ala	Val	Ser	Leu	Xaa
	35				40					45					
Arg	His	Xaa	Gln	Xaa	His	Glu	Val	Arg	Ala	Pro	Pro	Gly	Thr	Phe	Ala
50				55					60						65
Cys	Thr	Xaa	Cys	Gly	Gln	Asp	Phe	Ala	Gln	Glu	Xaa	Gly	Leu	His	Gln
			70						75					80	
His	Tyr	Ile	Arg	His	Ala	Arg	Gly	Gly	Leu						
			85				90								

<210> 415  
 <211> 190  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -82...-1

<400> 415

Met	Tyr	Val	Trp	Pro	Cys	Ala	Val	Val	Leu	Ala	Gln	Tyr	Leu	Trp	Phe
		-80					-75				-70				
His	Arg	Arg	Ser	Leu	Pro	Gly	Lys	Ala	Ile	Leu	Glu	Ile	Gly	Ala	Gly
	-65					-60					-55				
Val	Ser	Leu	Pro	Gly	Ile	Leu	Ala	Ala	Lys	Cys	Gly	Ala	Glu	Val	Ile
-50				-45						-40				-35	
Leu	Ser	Asp	Ser	Ser	Glu	Leu	Pro	His	Cys	Leu	Glu	Val	Cys	Arg	Gln
			-30					-25					-20		
Ser	Cys	Gln	Met	Asn	Asn	Leu	Pro	His	Leu	Gln	Val	Val	Gly	Leu	Thr



-15					-10					-5					
Trp	Gly	His	Ile	Ser	Trp	Asp	Leu	Leu	Ala	Leu	Pro	Pro	Gln	Asp	Ile
	1					5					10				
Ile	Leu	Ala	Ser	Asp	Val	Phe	Phe	Glu	Pro	Glu	Xaa	Phe	Glu	Asp	Ile
15					20					25					30
Leu	Ala	Thr	Ile	Tyr	Phe	Leu	Met	His	Lys	Asn	Pro	Lys	Val	Gln	Leu
					35					40					45
Trp	Ser	Thr	Tyr	Gln	Val	Arg	Xaa	Ala	Asp	Trp	Ser	Leu	Glu	Ala	Leu
					50					55					60
Leu	Tyr	Lys	Trp	Asp	Met	Lys	Cys	Val	His	Ile	Pro	Leu	Glu	Ser	Phe
					65					70					75
Asp	Ala	Asp	Lys	Glu	Xaa	Ile	Ala	Glu	Ser	Thr	Leu	Pro	Gly	Arg	His
					80					85					90
Thr	Val	Glu	Met	Leu	Val	Ile	Ser	Phe	Ala	Lys	Asp	Ser	Leu		
95					100					105					

<210> 416

<211> 114

<212> PRT

213> Homo sapiens

 $\langle 220 \rangle$ 

...<221> SIGNAL

 $\langle 222 \rangle - 60..-1$ 
$$400 > 416$$

Met Met Ala Ala Val Pro Pro Gly Leu Glu Pro Trp Asn Arg Val Arg  
60 -55 -50 -45

Ile Pro Lys Ala Gly Asn Arg Ser Ala Val Thr Val Gln Asn Pro Gly  
 -40 -35 -30

Ala Ala Leu Asp Leu Cys Ile Ala Ala Val Ile Lys Glu Cys His Leu  
-25 -20 -15

Val	Ile	Leu	Ser	Leu	Lys	Ser	Gln	Thr	Leu	Asp	Ala	Glu	Thr	Asp	Val
		-10					-5					1			

Leu Cys Ala Val Leu Tyr Ser Asn His Asn Arg Met Gly Arg His Lys  
5 10 15 20

Pro His Leu Ala Leu Lys Gln Val Glu Gln Cys Leu Lys Arg Leu Lys  
25 30 35

Asn Met Asn Leu Glu Gly Ser Ile Gln Asp Leu Phe Glu Leu Phe Ser  
40 45 50

Ser Lys

<210> 417

<211> 161

<212> PRT

<213> Homo sapiens

 $\langle 220 \rangle$ 

<221> SIGNAL

<222> -108..-1

<400> 417

Met Thr Ser Gly Gln Ala Arg Ala Ser Xaa Gln Ser Pro Gln Ala Leu

[illegible]

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210> 418
211> 67
212> PRT
213> Homo sapiens

```

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<220>
<221> SIGNAL
<222> -21...-1

```

```

400> 418
Met Leu Gly Gly Asp His Arg Ala Leu Leu Leu Lys Ile Trp Leu Leu
-20 -15 -10
Gln Arg Pro Glu Ser Gln Glu Gly Leu Leu Pro Gly Arg Leu Val Val
5 1 5 10
Met Glu Arg Arg Val Lys Asn Asp Leu Met Ser Phe Leu Ser Thr Val
15 20 25
Leu Leu Ser Phe His Ser Ser Asn Ala Arg Val Ser His Cys Glu Pro
30 35 40
Leu Arg Met
45

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```
<210> 419
<211> 332
<212> PRT
<213> Homo sapiens
```

```
<220>
<221> SIGNAL
<222> -32...-1
```

<400> 419  
Met Ile Xaa Leu Arg Asp Thr Ala Ala Ser Leu Arg Leu Glu Arg Asp

-30							-25					-20				
Thr	Arg	Gln	Leu	Pro	Leu	Leu	Thr	Ser	Ala	Leu	His	Gly	Leu	Gln	Gln	
	-15					-10					-5					
Gln	His	Pro	Ala	Phe	Ser	Gly	Val	Ala	Arg	Leu	Ala	Lys	Arg	Trp	Val	
1				5				10						15		
Arg	Ala	Gln	Leu	Leu	Gly	Glu	Gly	Phe	Ala	Asp	Glu	Ser	Leu	Asp	Leu	
			20					25					30			
Val	Ala	Ala	Ala	Leu	Phe	Leu	His	Pro	Glu	Pro	Phe	Thr	Pro	Pro	Ser	
		35				40						45				
Ser	Pro	Gln	Val	Gly	Phe	Leu	Arg	Phe	Leu	Phe	Leu	Val	Ser	Thr	Phe	
	50					55					60					
Asp	Trp	Lys	Asn	Asn	Pro	Leu	Phe	Val	Asn	Leu	Asn	Asn	Glu	Leu	Thr	
65					70					75					80	
Val	Glu	Glu	Gln	Val	Glu	Ile	Arg	Ser	Gly	Phe	Leu	Ala	Ala	Arg	Ala	
				85					90					95		
Gln	Leu	Pro	Val	Met	Val	Ile	Val	Thr	Pro	Gln	Xaa	Arg	Lys	Asn	Ser	
			100					105					110			
Val	Trp	Thr	Gln	Asp	Gly	Pro	Ser	Ala	Gln	Ile	Leu	Gln	Gln	Leu	Val	
		115				120						125				
Val	Leu	Ala	Ala	Glu	Xaa	Leu	Pro	Met	Leu	Xaa	Xaa	Gln	Leu	Met	Asp	
	130					135					140					
Pro	Arg	Gly	Pro	Gly	Asp	Ile	Arg	Thr	Xaa	Phe	Arg	Pro	Pro	Leu	Asp	
145					150					155					160	
Ile	Tyr	Asp	Val	Leu	Ile	Arg	Leu	Ser	Pro	Arg	His	Ile	Pro	Arg	His	
				165					170					175		
Arg	Gln	Ala	Val	Asp	Ser	Pro	Ala	Ala	Ser	Phe	Cys	Arg	Gly	Leu	Leu	
			180					185					190			
Ser	Gln	Pro	Gly	Pro	Ser	Ser	Leu	Met	Pro	Val	Leu	Gly	Xaa	Asp	Pro	
		195				200					205					
Pro	Gln	Leu	Tyr	Leu	Thr	Gln	Leu	Xaa	Glu	Ala	Phe	Gly	Asp	Leu	Ala	
	210					215					220					
Leu	Phe	Phe	Tyr	Asp	Gln	His	Gly	Gly	Glu	Val	Ile	Gly	Val	Leu	Trp	
225					230					235					240	
Lys	Pro	Thr	Ser	Phe	Gln	Pro	Gln	Pro	Phe	Lys	Ala	Ser	Ser	Thr	Lys	
				245					250					255		
Gly	Arg	Met	Val	Met	Ser	Arg	Gly	Gly	Glu	Leu	Val	Met	Val	Pro	Asn	
			260					265					270			
Val	Glu	Ala	Ile	Leu	Glu	Asp	Phe	Ala	Val	Leu	Gly	Glu	Gly	Leu	Val	
		275				280						285				
Gln	Thr	Val	Glu	Ala	Arg	Ser	Glu	Arg	Trp	Thr	Val					
						295					300					

```
<210> 420
<211> 65
<212> PRT
<213> Homo sapiens
```

```
<220>
<221> SIGNAL
<222> -19..-1
```

```

<400> 420
Met Gly Gly Ile Trp Asn Ala Leu Ser Met Ser Ser Phe Ser Phe His
      -15                      -10                      -5
Ser Ser Ser Cys Ser Ala Leu Ser Ala Lys Ser Leu Leu Ser Arg His

```

	1					5						10					
His	Ile	Leu	Gln	Gln	Phe	Leu	Val	Arg	Lys	Ser	Val	Pro	Leu	Glu	Asn		
	15					20					25						
Ala	Ser	Leu	Pro	Phe	Pro	His	Leu	Gly	Ser	Ser	Leu	Phe	Lys	Ile	Val		
30					35					40					45		
Gly																	

<210> 421  
 <211> 57  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -30..-1

Met	Pro	Thr	Gly	Lys	Gln	Leu	Ala	Asp	Ile	Gly	Tyr	Lys	Thr	Phe	Ser		
-30					-25					-20					-15		
Thr	Ser	Met	Met	Leu	Leu	Thr	Val	Tyr	Gly	Gly	Tyr	Leu	Cys	Ser	Val		
				-10					-5					1			
Arg	Val	Tyr	His	Tyr	Phe	Gln	Trp	Arg	Arg	Ala	Gln	Arg	Gln	Ala	Ala		
	5					10						15					
Glu	Glu	Gln	Lys	Xaa	Ser	Gly	Ile	Met									
20						25											

<210> 422  
 <211> 85  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -17..-1

Met	Lys	Lys	Val	Leu	Leu	Leu	Ile	Thr	Ala	Ile	Leu	Ala	Val	Ala	Val		
	-15						-10				-5						
Gly	Phe	Pro	Val	Ser	Gln	Asp	Gln	Glu	Arg	Glu	Lys	Arg	Ser	Ile	Ser		
1				5					10					15			
Asp	Ser	Asp	Glu	Leu	Ala	Ser	Gly	Xaa	Phe	Val	Phe	Pro	Tyr	Pro	Tyr		
			20				25						30				
Pro	Phe	Arg	Pro	Leu	Pro	Pro	Ile	Pro	Phe	Pro	Arg	Phe	Pro	Trp	Phe		
	35						40					45					
Arg	Arg	Asn	Phe	Pro	Ile	Pro	Ile	Pro	Glu	Ser	Ala	Pro	Thr	Thr	Pro		
	50					55						60					
Leu	Pro	Ser	Glu	Lys													
65																	

<210> 423  
 <211> 85

<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -17..-1

<400> 423  
Met Lys Lys Val Leu Leu Leu Ile Thr Ala Ile Leu Ala Val Ala Val  
          -15                  -10                  -5  
Gly Phe Pro Val Ser Gln Asp Xaa Glu Arg Glu Lys Arg Ser Ile Ser  
  1                  5                  10                  15  
Asp Ser Asp Glu Leu Ala Ser Gly Phe Phe Val Phe Pro Tyr Pro Tyr  
          20                  25                  30  
Pro Phe Arg Pro Leu Pro Pro Ile Pro Phe Pro Arg Phe Pro Trp Phe  
          35                  40                  45  
Arg Arg Asn Phe Pro Ile Pro Ile Pro Glu Ser Ala Pro Thr Thr Pro  
          50                  55                  60  
Leu Pro Ser Glu Lys  
          65

<210> 424  
<211> 69  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -29..-1

<400> 424  
Met Thr Cys Arg Gly Ser Cys Ser Tyr Ala Thr Arg Arg Ser Pro Ser  
                  -25                  -20                  -15  
Glu Leu Ser Leu Leu Pro Ser Ser Leu Trp Val Leu Ala Thr Ser Ser  
          -10                  -5                  1  
Pro Thr Ile Thr Ile Ala Leu Ala Met Ala Ala Gly Asn Leu Cys Pro  
  5                  10                  15  
Leu Pro Ser Ser Xaa Arg Xaa Lys Arg Arg Trp Cys Gln Ala Xaa Gln  
 20                  25                  30                  35  
Gln Xaa Ala Leu Leu  
          40

<210> 425  
<211> 122  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -56..-1

<400> 425  
Met Val Pro Trp Pro Arg Gly Lys Val Lys Thr Ala Pro Ile Pro Ile

```

-55          -50          -45
Ser Arg Phe Pro Phe Leu Pro Thr His Asp Pro Pro Thr Pro Ala His
-40          -35          -30          -25
Trp Ser Pro Ala Ser His Gln Gln Phe Lys His Xaa Ser Pro Leu Leu
          -20          -15          -10
Thr Leu Ala Leu Leu Gly Gln Cys Ser Leu Phe Xaa Asn Leu Arg Lys
          -5          1          5
Lys Leu Ala Gly Gln Lys Ala Lys Lys Leu Pro Ser Phe Ser Ser Leu
10          15          20
Pro Leu Thr Leu Trp Pro Leu Thr Pro Gln Phe Ala Glu Leu Thr Thr
25          30          35          40
Val Ala Gln Lys Lys Leu Arg Trp Ser Gly Thr Leu Gly Trp Gly Pro
          45          50          55
Val Pro Ser Trp Val Gln Phe Phe Leu Gly
          60          65

```

<210> 426  
 <211> 41  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -30...-1

```

<400> 426
Met Ala Cys Glu Thr His Gly Val Leu Val Pro Ala His Leu Ser Gly
30          -25          -20          -15
Leu Ile Thr Cys Leu Leu Ala Phe Trp Val Pro Ala Ser Cys Ile Gln
          -10          -5          1
Arg Cys Ser Gly Ser Pro Leu Pro Leu
5          10

```

<210> 427  
 <211> 50  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -36...-1

```

<400> 427
Met Ala Pro His Thr Ala Ser Phe Gly Val Cys Pro Leu Leu Ser Val
-35          -30          -25
Thr Arg Val Val Ala Thr Glu His Trp Leu Phe Leu Ala Ser Leu Ser
-20          -15          -10          -5
Gly Ile Lys Thr Tyr Gln Ser Tyr Ile Ser Val Phe Cys Lys Val Thr
1          5          10
Leu Ile

```

<210> 428  
 <211> 136  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -18..-1

<400> 428  
 Met Asp Ser Leu Arg Lys Met Leu Ile Ser Val Ala Met Leu Gly Ala  
                   -15                  -10                  -5  
 Xaa Ala Gly Val Gly Tyr Ala Leu Leu Val Ile Val Thr Pro Gly Glu  
           1                  5                  10  
 Arg Arg Lys Gln Glu Met Leu Lys Glu Met Pro Leu Gln Asp Pro Arg  
 15                  20                  25                  30  
 Ser Arg Glu Glu Ala Ala Arg Thr Gln Gln Leu Leu Leu Ala Thr Leu  
                   35                  40                  45  
 Gln Glu Ala Ala Thr Thr Gln Glu Asn Val Ala Trp Arg Lys Asn Trp  
                   50                  55                  60  
 Met Val Gly Gly Glu Gly Gly Ala Thr Gly Xaa His Arg Glu Thr Gly  
           65                  70                  75  
 Leu Ala Ser Val Gly Ala Gly Pro Trp Leu Gly Arg Arg Asn Pro Arg  
           80                  85                  90  
 Gln Leu Ser Pro Ser Trp Ala Xaa Arg Lys Ile Arg Xaa Glu Asn Xaa  
 95                  100                  105                  110  
 Met Pro Gly Leu Ser Gly Val Leu  
                   115

<210> 429  
 <211> 194  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -65..-1

<400> 429  
 Met Gln Asp Ala Pro Leu Ser Cys Leu Ser Pro Thr Lys Trp Ser Ser  
 -65                  -60                  -55                  -50  
 Val Ser Ser Ala Asp Ser Thr Glu Lys Ser Ala Ser Ala Ala Gly Thr  
                   -45                  -40                  -35  
 Arg Asn Leu Pro Phe Gln Phe Cys Leu Arg Gln Ala Leu Arg Met Lys  
                   -30                  -25                  -20  
 Ala Ala Gly Ile Leu Thr Leu Ile Gly Cys Leu Val Thr Gly Val Glu  
           -15                  -10                  -5  
 Ser Lys Ile Tyr Thr Arg Cys Lys Leu Ala Lys Ile Phe Ser Arg Ala  
   1                  5                  10                  15  
 Gly Leu Asp Asn Xaa Arg Gly Phe Ser Leu Gly Asn Trp Ile Cys Met  
                   20                  25                  30  
 Ala Tyr Tyr Glu Ser Gly Tyr Asn Thr Ala Gln Thr Val Leu Asp  
                   35                  40                  45  
 Asp Gly Ser Ile Asp Tyr Gly Ile Phe Gln Ile Asn Ser Phe Ala Trp  
   50                  55                  60

Cys Arg Arg Gly Lys Leu Lys Glu Asn Asn His Cys His Val Ala Cys  
65 70 75  
Ser Ala Leu Xaa Thr Asp Asp Leu Thr Asp Ala Ile Ile Cys Ala Xaa  
80 85 90 95  
Lys Ile Val Lys Glu Thr Gln Gly Met Asn Tyr Trp Gln Gly Trp Lys  
100 105 110  
Lys His Cys Glu Gly Arg Asp Leu Ser Xaa Trp Lys Lys Gly Cys Glu  
115 120 125  
Val Ser

<210> 430  
<211> 141  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -69..-1

<400> 430  
Met Thr Ser Gln Pro Val Pro Asn Glu Thr Ile Ile Val Leu Pro Ser  
-65 -60 -55  
Asn Val Ile Asn Phe Ser Gln Ala Glu Lys Pro Glu Pro Thr Asn Gln  
-50 -45 -40  
Gly Gln Asp Ser Leu Lys Lys His Leu His Ala Glu Ile Lys Val Ile  
-35 -30 -25  
Gly Thr Ile Gln Ile Leu Cys Gly Met Met Val Leu Ser Leu Gly Ile  
-20 -15 -10  
Ile Leu Ala Ser Ala Ser Phe Ser Pro Asn Phe Thr Gln Val Thr Ser  
-5 1 5 10  
Thr Leu Leu Asn Ser Ala Tyr Pro Phe Ile Gly Pro Phe Phe Val Xaa  
15 20 25  
Lys Xaa Ser Glu Glu Gly Arg Met Gly Gln Xaa Gly Glu Glu Xaa Xaa  
30 35 40  
Asn Ser Leu Asn Phe Pro Xaa Ala Ser Leu Leu Xaa Leu Ile Cys Gln  
45 50 55  
Xaa Gln Gly Phe Asn Gly Glu Ser Cys Ser Pro Val Gly  
60 65 70

<210> 431  
<211> 248  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -69..-1

<400> 431  
Met Thr Ser Gln Pro Val Pro Asn Glu Thr Ile Ile Val Leu Pro Ser  
-65 -60 -55  
Asn Val Ile Asn Phe Ser Gln Ala Glu Lys Pro Glu Pro Thr Asn Gln  
-50 -45 -40



Gly Gln Asp Ser Leu Lys Lys His Leu His Ala Glu Xaa Lys Val Ile  
 -35 -30 -25  
 Gly Thr Ile Gln Ile Leu Cys Gly Met Met Val Leu Ser Leu Gly Ile  
 -20 -15 -10  
 Ile Leu Ala Ser Ala Ser Phe Ser Pro Asn Phe Thr Gln Val Thr Ser  
 -5 1 5 10  
 Thr Leu Leu Asn Ser Ala Tyr Pro Phe Ile Gly Pro Phe Phe Phe Ile  
 15 20 25  
 Ile Ser Gly Ser Leu Ser Ile Ala Thr Lys Lys Arg Leu Thr Asn Leu  
 30 35 40  
 Leu Val His Thr Thr Leu Val Gly Ser Ile Leu Ser Ala Leu Ser Ala  
 45 50 55  
 Leu Val Gly Phe Ile Xaa Leu Ser Val Lys Gln Ala Thr Leu Asn Pro  
 60 65 70 75  
 Ala Ser Leu Xaa Cys Glu Leu Xaa Lys Asn Asn Ile Pro Thr Xaa Xaa  
 80 85 90  
 Tyr Val Xaa Tyr Phe Tyr His Asp Ser Leu Tyr Thr Thr Asp Xaa Tyr  
 95 100 105  
 Thr Ala Lys Ala Xaa Leu Ala Gly Thr Leu Ser Leu Met Leu Ile Cys  
 110 115 120  
 Thr Leu Leu Glu Phe Cys Xaa Xaa Val Leu Thr Ala Val Leu Arg Trp  
 125 130 135  
 Lys Gln Ala Tyr Ser Asp Phe Pro Gly Ser Val Leu Phe Leu Pro Xaa  
 140 145 150 155  
 Ser Tyr Ile Gly Asn Ser Gly Met Ser Ser Lys Met Thr His Asp Cys  
 160 165 170  
 Gly Tyr Glu Glu Leu Leu Thr Ser  
 175

<210> 432  
 <211> 49  
 <212> PRT  
 <213> Homo sapiens  
 <220>  
 <221> SIGNAL  
 <222> -36...-1

<400> 432  
 Met Gln Val Pro His Leu Arg Val Trp Thr Gln Val Xaa Asp Thr Phe  
 -35 -30 -25  
 Ile Gly Tyr Arg Asn Leu Gly Phe Thr Ser Met Cys Ile Leu Phe His  
 -20 -15 -10 -5  
 Cys Leu Leu Ser Phe Gln Val Phe Lys Lys Lys Arg Lys Leu Xaa Leu  
 1 5 10  
 Phe

<210> 433  
 <211> 86  
 <212> PRT  
 <213> Homo sapiens  
 <220>

<221> SIGNAL  
<222> -14..-1

<400> 433

Met	Val	Ala	Leu	Asn	Leu	Ile	Leu	Val	Pro	Cys	Cys	Ala	Ala	Trp	Cys	
			-10						-5					1		
Asp	Pro	Arg	Arg	Ile	His	Ser	Gln	Asp	Asp	Val	Leu	Arg	Ser	Ser	Ala	
	5						10					15				
Ala	Asp	Thr	Gly	Ser	Ala	Met	Gln	Arg	Arg	Glu	Ala	Trp	Ala	Gly	Trp	
	20					25					30					
Arg	Arg	Ser	Gln	Pro	Phe	Ser	Val	Gly	Leu	Pro	Ser	Ala	Glu	Arg	Leu	
35				40					45						50	
Glu	Asn	Gln	Pro	Gly	Lys	Leu	Ser	Trp	Arg	Ser	Leu	Val	Gly	Glu	Gly	
			55					60						65		
His	Arg	Ile	Cys	Asp	Leu											
			70													

<210> 434  
<211> 144  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -58..-1

<400> 434

Met	Thr	Arg	Leu	Cys	Leu	Pro	Arg	Pro	Glu	Ala	Arg	Glu	Asp	Pro	Ile	
			-55					-50					-45			
Pro	Val	Pro	Pro	Arg	Gly	Leu	Gly	Ala	Gly	Glu	Gly	Ser	Gly	Ser	Pro	
		-40					-35					-30				
Val	Arg	Pro	Pro	Val	Ser	Thr	Trp	Gly	Pro	Ser	Trp	Ala	Gln	Leu	Leu	
		-25				-20					-15					
Asp	Ser	Val	Leu	Trp	Leu	Gly	Ala	Leu	Gly	Leu	Thr	Ile	Gln	Ala	Val	
10				-5					1					5		
Phe	Ser	Thr	Thr	Gly	Pro	Ala	Leu	Leu	Leu	Leu	Val	Ser	Phe	Leu		
		10				15					20					
Thr	Phe	Asp	Leu	Leu	His	Arg	Pro	Ala	Val	Thr	Leu	Cys	His	Ser	Ala	
		25				30					35					
Asn	Phe	Ser	Pro	Gly	Ala	Arg	Val	Arg	Gly	Pro	Val	Lys	Val	Leu	Asp	
	40				45					50						
Ser	Arg	Arg	Leu	Tyr	Ser	Cys	Lys	Trp	Val	Gln	Ser	Gln	Asp	Asn	Leu	
55				60					65					70		
Ala	Ser	Arg	Lys	His	Cys	Cys	Cys	Cys	Ser	Trp	Gly	Trp	Ala	Arg	Ser	
			75					80								

<210> 435  
<211> 121  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL

<222> -16..-1

<400> 435

Met	Glu	Arg	Leu	Val	Leu	Thr	Leu	Cys	Thr	Leu	Pro	Leu	Ala	Val	Ala
	-15					-10					-5				
Ser	Ala	Gly	Cys	Ala	Thr	Thr	Pro	Ala	Arg	Asn	Leu	Ser	Cys	Tyr	Gln
1				5					10					15	
Cys	Phe	Lys	Val	Ser	Ser	Trp	Thr	Glu	Cys	Pro	Pro	Thr	Trp	Cys	Ser
			20					25					30		
Pro	Leu	Asp	Gln	Val	Cys	Ile	Ser	Asn	Glu	Val	Val	Val	Ser	Phe	Ser
		35				40						45			
Glu	Ser	Pro	Pro	Gly	Arg	Gly	Xaa	Val	Pro	Xaa	Ala	Gly	Glu	Xaa	Pro
	50					55					60				
Val	Pro	Pro	Pro	Leu	Xaa	Asp	Leu	Xaa	Met	Thr	Pro	Arg	Xaa	Xaa	Arg
65					70					75					80
Ala	Trp	Gly	Pro	Val	Gly	Pro	Lys	Val	Pro	Pro	Ala	Val	Ser	Pro	Ala
				85					90					95	
Leu	Gly	Ser	Gly	Glu	His	Pro	Xaa	Xaa							
			100					105							

$\langle 210 \rangle$  436

$\angle 211 > 162$

23<212> PRT

<213> Homo sapiens

 $\langle 220 \rangle$ 

<221> SIGNAL

 $\langle 222 \rangle \quad -16 \dots -1$ 

<400> 436

[illegible]

<210> 437  
<211> 110  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -20...-1

<400> 437  
Met Xaa Leu Met Val Leu Val Phe Thr Ile Gly Leu Thr Leu Leu Leu  
-20 -15 -10 -5  
Gly Xaa Gln Ala Met Pro Ala Asn Arg Leu Ser Cys Tyr Arg Lys Ile  
1 5 10  
Leu Lys Asp His Asn Cys His Asn Leu Pro Glu Gly Val Ala Asp Leu  
15 20 25  
Thr Gln Ile Asp Val Asn Val Gln Asp His Phe Trp Asp Gly Lys Gly  
30 35 40  
Cys Glu Met Ile Cys Tyr Cys Asn Phe Lys Arg Ile Ala Leu Leu Pro  
45 50 55 60  
Lys Arg Arg Phe Leu Trp Thr Lys Asp Leu Phe Arg Asp Ser Leu Gln  
65 70 75  
Gln Ser Met Arg Ile Phe Met Tyr Ser Gly Glu His His Ser  
80 85 90

<210> 438  
<211> 71  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -15...-1

<400> 438  
Met Lys Leu Leu Thr His Asn Leu Leu Ser Ser His Val Arg Gly Val  
-15 -10 -5 1  
Gly Ser Arg Gly Phe Pro Leu Arg Leu Gln Ala Thr Glu Val Arg Ile  
5 10 15  
Cys Pro Val Glu Phe Asn Pro Asn Phe Val Ala Arg Met Ile Pro Lys  
20 25 30  
Val Glu Trp Ser Ala Phe Leu Glu Ala Xaa Asp Asn Leu Arg Leu Ile  
35 40 45  
Gln Val Pro Arg Arg Ala Gly  
50 55

<210> 439  
<211> 99  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL

<222> -24..-1

<400> 439

Met	Lys	Ser	Ala	Lys	Leu	Gly	Phe	Leu	Leu	Arg	Phe	Phe	Ile	Phe	Cys
				-20					-15					-10	
Ser	Leu	Asn	Thr	Leu	Leu	Leu	Gly	Gly	Val	Asn	Lys	Ile	Ala	Glu	Lys
			-5				1				5				
Ile	Cys	Gly	Asp	Leu	Lys	Asp	Pro	Cys	Lys	Leu	Asp	Met	Asn	Phe	Gly
10					15					20					
Ser	Cys	Tyr	Glu	Val	His	Phe	Arg	Tyr	Phe	Tyr	Asn	Arg	Thr	Ser	Lys
25					30				35					40	
Arg	Cys	Glu	Thr	Phe	Val	Phe	Ser	Ser	Cys	Asn	Gly	Asn	Leu	Asn	Asn
				45					50					55	
Phe	Lys	Leu	Lys	Ile	Glu	Arg	Glu	Val	Xaa	Cys	Val	Ala	Lys	Tyr	Lys
			60					65						70	
Pro	Pro	Arg													
															75

<210> 440

<211> 169

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -25..-1

<400> 440

Met	Arg	Lys	Pro	Ala	Ala	Gly	Phe	Leu	Pro	Ser	Leu	Leu	Lys	Val	Leu
25				-20					-15					-10	
Leu	Leu	Pro	Leu	Ala	Pro	Ala	Ala	Ala	Gln	Asp	Ser	Thr	Gln	Ala	Ser
				-5				1				5			
Thr	Pro	Gly	Ser	Pro	Leu	Ser	Pro	Thr	Glu	Tyr	Gln	Arg	Phe	Phe	Ala
10						15					20				
Leu	Leu	Thr	Pro	Thr	Trp	Lys	Ala	Glu	Thr	Thr	Cys	Arg	Leu	Arg	Ala
25					30					35					
Thr	His	Gly	Cys	Arg	Asn	Pro	Thr	Leu	Val	Gln	Leu	Asp	Gln	Tyr	Glu
40					45				50					55	
Asn	His	Gly	Leu	Val	Pro	Asp	Gly	Ala	Val	Cys	Ser	Asn	Leu	Pro	Tyr
				60					65					70	
Ala	Ser	Trp	Phe	Glu	Ser	Phe	Cys	Gln	Phe	Thr	His	Tyr	Arg	Cys	Ser
			75					80					85		
Asn	His	Val	Tyr	Tyr	Ala	Lys	Arg	Val	Leu	Cys	Ser	Gln	Pro	Val	Ser
		90				95						100			
Ile	Leu	Ser	Pro	Asn	Thr	Leu	Lys	Glu	Ile	Glu	Xaa	Ser	Ala	Glu	Val
105					110					115					
Ser	Pro	Thr	Thr	Asp	Asp	Leu	Pro	His	Leu	Thr	Pro	Leu	His	Ser	Asp
120				125						130					135
Arg	Thr	Pro	Asp	Leu	Pro	Ala	Leu	Ala							
															140

<210> 441

<211> 167

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -76..-1

<400> 441

```
Met Gly Asp Tyr Leu Leu Arg Gly Tyr Arg Met Leu Gly Glu Thr Cys
  -75                      -70                      -65
Ala Asp Cys Gly Thr Ile Leu Leu Gln Asp Lys Gln Arg Lys Ile Tyr
-60                      -55                      -50                      -45
Cys Val Ala Cys Gln Glu Leu Asp Ser Asp Val Asp Lys Asp Asn Pro
                      -40                      -35                      -30
Ala Leu Asn Ala Gln Ala Ala Leu Ser Gln Ala Arg Glu His Gln Leu
                      -25                      -20                      -15
Ala Ser Ala Ser Glu Leu Pro Leu Gly Ser Arg Pro Ala Pro Gln Pro
                      -10                      -5                      1
Pro Val Pro Arg Pro Glu His Cys Glu Gly Ala Ala Ala Gly Leu Lys
5                      10                      15                      20
Ala Ala Gln Gly Pro Pro Ala Pro Ala Val Pro Pro Asn Thr Xaa Val
                      25                      30                      35
Met Ala Cys Thr Gln Thr Ala Leu Leu Gln Lys Leu Thr Trp Ala Ser
                      40                      45                      50
Ala Glu Leu Gly Ser Xaa Thr Ser Xaa Gly Lys Xaa Ala Ser Ser Cys
                      55                      60                      65
Val Ala Leu Ser Ala His Val Arg Arg Pro Cys Ala Ala Cys Ser Ser
                      70                      75                      80
Tyr Ser Thr Lys Arg Ser Pro
85                      90
```

<210> 442

<211> 70

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -15..-1

<400> 442

```
Met Ile Leu Cys Phe Leu Leu Pro His His Arg Leu Gln Glu Ala Arg
-15                      -10                      -5                      1
Gln Ile Gln Val Leu Lys Met Leu Pro Arg Glu Lys Leu Arg Arg Arg
                      5                      10                      15
Glu Glu Arg Lys Gln Ile Asn Gly Lys Lys Xaa Arg Thr Lys Tyr Glu
                      20                      25                      30
Thr Pro Arg Lys Xaa Xaa Gly Lys Lys Gly Gly Asn Xaa Xaa Xaa Xaa
35                      40                      45
Xaa Leu Ser Lys Arg Asp
50                      55
```

<210> 443

<211> 381  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -33..-1

<400> 443

Met	Ser	Trp	Thr	Val	Pro	Val	Val	Arg	Ala	Ser	Gln	Arg	Val	Ser	Ser	
			-30					-25					-20			
Val	Gly	Ala	Asn	Xaa	Leu	Cys	Leu	Gly	Met	Ala	Leu	Cys	Pro	Arg	Gln	
	-15						-10					-5				
Ala	Thr	Arg	Ile	Pro	Leu	Asn	Gly	Thr	Trp	Leu	Phe	Thr	Pro	Val	Ser	
1				5					10						15	
Lys	Met	Ala	Thr	Val	Lys	Ser	Glu	Leu	Ile	Glu	Arg	Phe	Thr	Ser	Glu	
			20					25						30		
Lys	Pro	Val	His	His	Ser	Lys	Val	Ser	Ile	Ile	Gly	Thr	Gly	Ser	Val	
			35					40					45			
Gly	Met	Ala	Cys	Ala	Ile	Ser	Ile	Leu	Leu	Lys	Gly	Leu	Ser	Asp	Glu	
	50						55					60				
Leu	Ala	Leu	Val	Asp	Leu	Asp	Glu	Xaa	Lys	Leu	Lys	Gly	Glu	Thr	Met	
65					70						75					
Asp	Leu	Gln	His	Gly	Ser	Pro	Phe	Thr	Lys	Met	Pro	Asn	Ile	Val	Cys	
80					85				90						95	
Ser	Lys	Xaa	Tyr	Phe	Val	Thr	Ala	Asn	Ser	Asn	Leu	Val	Ile	Ile	Thr	
			100					105						110		
Ala	Gly	Ala	Arg	Gln	Xaa	Lys	Gly	Glu	Thr	Arg	Leu	Asn	Leu	Xaa	Gln	
			115					120					125			
Arg	Asn	Val	Ala	Ile	Phe	Lys	Leu	Met	Ile	Ser	Ser	Ile	Val	Gln	Tyr	
	130						135						140			
Ser	Pro	His	Cys	Lys	Leu	Ile	Ile	Val	Ser	Asn	Pro	Val	Asp	Ile	Leu	
	145				150					155						
Thr	Tyr	Val	Ala	Trp	Lys	Leu	Ser	Ala	Phe	Pro	Lys	Asn	Arg	Ile	Ile	
160					165				170						175	
Gly	Ser	Gly	Cys	Asn	Leu	Ile	Xaa	Ala	Arg	Phe	Arg	Phe	Leu	Ile	Gly	
			180						185					190		
Gln	Lys	Leu	Gly	Ile	His	Ser	Glu	Ser	Cys	His	Gly	Trp	Ile	Leu	Gly	
			195					200					205			
Glu	His	Gly	Asp	Ser	Ser	Val	Pro	Val	Trp	Ser	Gly	Val	Asn	Ile	Ala	
	210						215					220				
Gly	Val	Pro	Leu	Lys	Asp	Leu	Asn	Ser	Asp	Ile	Gly	Thr	Asp	Lys	Asp	
	225					230					235					
Pro	Glu	Gln	Trp	Lys	Asn	Val	His	Lys	Glu	Val	Thr	Ala	Thr	Ala	Tyr	
240					245					250					255	
Glu	Ile	Ile	Lys	Met	Lys	Gly	Tyr	Thr	Ser	Trp	Ala	Ile	Gly	Leu	Ser	
			260						265					270		
Val	Ala	Asp	Leu	Thr	Glu	Ser	Ile	Leu	Lys	Asn	Leu	Arg	Arg	Ile	His	
			275					280					285			
Pro	Val	Ser	Thr	Ile	Thr	Lys	Gly	Leu	Tyr	Gly	Ile	Xaa	Glu	Glu	Val	
			290				295					300				
Phe	Leu	Ser	Ile	Pro	Cys	Ile	Leu	Gly	Glu	Asn	Gly	Ile	Thr	Asn	Leu	
	305					310					315					
Ile	Lys	Ile	Lys	Leu	Thr	Pro	Glu	Glu	Glu	Ala	His	Leu	Lys	Lys	Ser	
320					325					330					335	
Ala	Lys	Thr	Leu	Trp	Glu	Ile	Gln	Asn	Lys	Leu	Lys	Leu				
			340						345							

<210> 444  
<211> 39  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -14..-1

<400> 444  
Met Tyr Tyr Met Val Cys Leu Phe Phe Arg Leu Ile Phe Ser Glu His  
                  -10                  -5                  1  
Leu Pro Ile Ile Gly Thr Val Thr Ser His Lys Thr Gly Thr Leu Thr  
      5                  10                  15  
Val Tyr Pro Thr Ser Ala Gly  
      20                  25

<210> 445  
<211> 50  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -37..-1

<400> 445  
Met Val Leu Thr Thr Leu Pro Leu Pro Ser Ala Asn Ser Pro Val Asn  
          -35                  -30                  -25  
Met Pro Thr Thr Gly Pro Asn Ser Leu Ser Tyr Ala Ser Ser Ala Leu  
      -20                  -15                  -10  
Ser Pro Cys Leu Thr Ala Pro Lys Ser Pro Arg Leu Ala Met Met Pro  
      5                  1                  5                  10  
Asp Asn

<210> 446  
<211> 51  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -26..-1

<400> 446  
Met Thr Pro Trp Cys Leu Ala Cys Leu Gly Arg Arg Pro Leu Ala Ser  
      -25                  -20                  -15  
Leu Gln Trp Ser Leu Thr Leu Ala Trp Cys Gly Ser Gly Ser His Trp  
      -10                  -5                  1                  5  
Thr Glu Arg Pro Xaa Gln Xaa Ser Pro Trp Xaa Ser Leu Ser Ala Thr



10 15 20  
Thr Arg Gly  
25

<210> 447  
<211> 242  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -30..-1

<400> 447  
Met Gly Glu Ala Ser Pro Pro Ala Pro Ala Arg Arg His Leu Leu Val  
-30 -25 -20 -15  
Leu Leu Leu Leu Leu Ser Thr Leu Val Ile Pro Ser Ala Ala Ala Pro  
-10 -5 1  
Ile His Asp Ala Asp Ala Gln Glu Ser Ser Leu Gly Leu Thr Gly Leu  
5 10 15  
Gln Ser Leu Leu Gln Gly Phe Ser Arg Leu Phe Leu Lys Gly Asn Leu  
20 25 30  
Leu Arg Gly Ile Asp Ser Leu Phe Ser Ala Pro Met Asp Phe Arg Gly  
35 40 45 50  
Leu Pro Gly Asn Tyr His Lys Glu Glu Asn Gln Glu His Gln Leu Gly  
55 60 65  
Asn Asn Thr Leu Ser Ser His Leu Gln Ile Asp Lys Met Thr Asp Asn  
70 75 80  
Lys Thr Gly Glu Val Leu Ile Ser Glu Asn Val Val Ala Ser Ile Gln  
85 90 95  
Pro Xaa Glu Gly Xaa Phe Glu Gly Asp Leu Lys Val Pro Arg Met Glu  
100 105 110  
Glu Lys Glu Ala Leu Val Pro Xaa Gln Lys Ala Thr Asp Ser Phe His  
115 120 125 130  
Thr Glu Leu His Pro Arg Val Ala Phe Trp Ile Ile Lys Leu Pro Arg  
135 140 145  
Arg Arg Ser His Gln Asp Ala Leu Glu Gly Gly His Trp Leu Xaa Glu  
150 155 160  
Lys Arg His Arg Leu Gln Ala Ile Arg Asp Gly Leu Arg Lys Gly Thr  
165 170 175  
His Lys Asp Xaa Leu Xaa Xaa Gly Thr Glu Ser Ser Ser His Ser Arg  
180 185 190  
Leu Ser Pro Arg Lys Xaa His Leu Leu Tyr Ile Leu Xaa Pro Ser Arg  
195 200 205 210  
Gln Leu

<210> 448  
<211> 154  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL

<222> -60..-1

<400> 448

Met	Gly	Ser	Lys	Cys	Cys	Lys	Gly	Gly	Pro	Asp	Glu	Asp	Ala	Val	Glu
-60				-55					-50						-45
Arg	Gln	Arg	Arg	Gln	Lys	Leu	Leu	Leu	Ala	Gln	Leu	His	His	Arg	Lys
			-40					-35						-30	
Arg	Val	Lys	Ala	Ala	Gly	Gln	Ile	Gln	Ala	Trp	Trp	Arg	Gly	Val	Leu
		-25				-20						-15			
Val	Arg	Arg	Thr	Leu	Leu	Val	Ala	Ala	Leu	Arg	Ala	Trp	Met	Ile	Gln
	-10				-5						1				
Cys	Trp	Trp	Arg	Thr	Leu	Val	Gln	Arg	Arg	Ile	Arg	Gln	Arg	Arg	Gln
5			10					15						20	
Ala	Leu	Leu	Gly	Val	Tyr	Val	Ile	Gln	Glu	Gln	Ala	Ala	Val	Lys	Leu
		25						30					35		
Gln	Ser	Cys	Ile	Arg	Met	Trp	Gln	Cys	Arg	Gln	Cys	Tyr	Arg	Gln	Met
		40				45						50			
Cys	Asn	Ala	Leu	Cys	Leu	Phe	Gln	Val	Pro	Lys	Ser	Ser	Leu	Ala	Phe
	55					60					65				
Gln	Thr	Asp	Gly	Phe	Leu	Gln	Val	Gln	Tyr	Ala	Ile	Pro	Ser	Lys	Gln
	70				75			80							
Pro	Glu	Phe	His	Ile	Glu	Ile	Leu	Ser	Ile						
85					90										

<210> 449

<211> 89

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -61..-1

<400> 449

Met	Asn	Ala	Ala	Ile	Asn	Thr	Gly	Pro	Ala	Pro	Ala	Val	Thr	Lys	Thr
-60					-55					-50					
Glu	Thr	Glu	Val	Gln	Asn	Pro	Asp	Val	Leu	Trp	Asp	Leu	Asp	Ile	Pro
-45				-40				-35						-30	
Glu	Ala	Arg	Ser	His	Ala	Asp	Gln	Asp	Ser	Asn	Pro	Lys	Ala	Glu	Ala
			-25				-20					-15			
Leu	Leu	Pro	Cys	Asn	Leu	His	Cys	Ser	Trp	Leu	His	Ser	Ser	Pro	Arg
		-10				-5					1				
Pro	Asp	Pro	His	Ser	His	Phe	Pro	Ser	Xaa	Arg	Arg	Cys	Pro	Leu	Pro
5					10					15					
His	Pro	Cys	Ala	Thr	Tyr	Pro	Pro	Xaa							
20				25											

<210> 450

<211> 73

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL  
<222> -26..-1

<400> 450

Met	Arg	Met	Ser	Leu	Ala	Gln	Arg	Val	Leu	Leu	Thr	Trp	Leu	Phe	Thr
-25					-20						-15				
Leu	Leu	Phe	Leu	Ile	Met	Leu	Val	Leu	Lys	Leu	Asp	Glu	Lys	Ala	Pro
-10				-5				1						5	
Trp	Asn	Trp	Phe	Leu	Ile	Phe	Ile	Pro	Val	Trp	Ile	Phe	Asp	Thr	Ile
			10				15						20		
Leu	Leu	Val	Leu	Leu	Ile	Val	Lys	Met	Ala	Gly	Arg	Cys	Lys	Ser	Gly
		25				30						35			
Phe	Asp	Leu	Asp	Met	Asp	His	Thr	Ile							
40						45									

<210> 451  
<211> 54  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -34..-1

<400> 451

Met	Ile	Pro	Leu	Ile	Ser	His	Leu	Ala	Glu	Ala	Ala	Pro	Pro	Thr	Ser
			-30					-25						-20	
Trp	Ser	Leu	Ile	Ser	Ser	Val	Leu	Asn	Val	Gly	His	Leu	Leu	Phe	Ser
			-15				-10							-5	
Ser	Ala	Cys	Ser	Val	Ser	Leu	Glu	Ala	Leu	Ser	Thr	Arg	Asn	Ile	Lys
		1			5						10				
Ala	Ile	Ile	Leu	Met	Lys										
15				20											

<210> 452  
<211> 121  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -38..-1

<400> 452

Met	Glu	Ser	Pro	Gln	Leu	His	Cys	Ile	Leu	Asn	Ser	Asn	Ser	Val	Ala
		-35					-30						-25		
Cys	Ser	Phe	Ala	Val	Gly	Ala	Gly	Phe	Leu	Ala	Phe	Leu	Ser	Cys	Leu
		-20				-15						-10			
Ala	Phe	Leu	Val	Leu	Asp	Thr	Gln	Glu	Thr	Arg	Ile	Ala	Gly	Thr	Arg
	-5				1				5					10	
Phe	Lys	Thr	Ala	Phe	Gln	Leu	Leu	Asp	Phe	Ile	Leu	Ala	Val	Leu	Trp
			15				20						25		
Ala	Val	Val	Trp	Phe	Met	Gly	Phe	Cys	Phe	Leu	Ala	Asn	Gln	Trp	Gln

	30		35		40										
His	Ser	Pro	Pro	Lys	Glu	Xaa	Leu	Leu	Gly	Ser	Ser	Ser	Ala	Gln	Ala
	45						50					55			
Ala	Ile	Gly	Xaa	His	Leu	Leu	Leu	His	Pro	Cys	Leu	Asp	Ile	Pro	Xaa
	60					65					70				
Leu	Pro	Gly	Xaa	Pro	Gly	Pro	Pro	Lys							
75					80										

<210> 453  
 <211> 166  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -37..-1

	400	453													
Met	Ser	Thr	Val	Gly	Leu	Phe	His	Phe	Pro	Thr	Pro	Leu	Thr	Arg	Ile
		-35					-30					-25			
Cys	Pro	Ala	Pro	Trp	Gly	Leu	Arg	Leu	Trp	Glu	Lys	Leu	Thr	Leu	Leu
		-20				-15					-10				
Ser	Pro	Gly	Ile	Ala	Val	Thr	Pro	Val	Gln	Met	Ala	Gly	Lys	Lys	Asp
	5				1				5					10	
Tyr	Pro	Ala	Leu	Leu	Ser	Leu	Asp	Glu	Asn	Glu	Leu	Glu	Glu	Gln	Phe
		15						20					25		
Val	Lys	Gly	His	Gly	Pro	Gly	Gly	Gln	Ala	Thr	Asn	Lys	Thr	Ser	Asn
		30					35					40			
Cys	Val	Val	Leu	Lys	Xaa	Ile	Pro	Ser	Gly	Ile	Val	Val	Lys	Cys	His
	45				50					55					
Gln	Thr	Arg	Ser	Val	Asp	Gln	Asn	Arg	Lys	Leu	Ala	Arg	Lys	Ile	Leu
	60				65					70					75
Gln	Glu	Lys	Val	Xaa	Val	Phe	Tyr	Asn	Gly	Glu	Asn	Ser	Pro	Val	His
			80						85					90	
Lys	Glu	Lys	Arg	Glu	Ala	Ala	Lys	Lys	Lys	Gln	Glu	Arg	Lys	Lys	Arg
		95						100					105		
Ala	Lys	Glu	Thr	Leu	Glu	Lys	Lys	Xaa	Leu	Leu	Lys	Xaa	Leu	Trp	Glu
		110					115						120		
Ser	Ser	Lys	Lys	Val	His										
		125													

<210> 454  
 <211> 180  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -26..-1

	400	454													
Met	Gly	Ile	Gln	Thr	Ser	Pro	Val	Leu	Leu	Ala	Ser	Leu	Gly	Val	Gly
	-25						-20					-15			

```

Leu Val Thr Leu Leu Gly Leu Ala Val Gly Ser Tyr Leu Val Arg Arg
-10          -5          1          5
Ser Arg Arg Pro Gln Val Thr Leu Leu Asp Pro Asn Glu Lys Tyr Leu
          10          15          20
Leu Arg Leu Leu Asp Lys Thr Thr Val Ser His Asn Thr Lys Arg Phe
          25          30          35
Arg Phe Ala Leu Pro Thr Ala His His Thr Leu Gly Leu Pro Val Gly
          40          45          50
Lys His Ile Tyr Leu Ser Thr Arg Ile Asp Gly Ser Leu Val Ile Arg
55          60          65          70
Pro Tyr Thr Pro Val Thr Ser Asp Glu Asp Gln Gly Tyr Val Asp Leu
          75          80          85
Val Xaa Lys Val Tyr Leu Lys Gly Val His Pro Lys Phe Pro Glu Gly
          90          95          100
Gly Lys Met Ser Xaa Tyr Leu Asp Xaa Leu Lys Val Gly Asp Xaa Val
          105          110          115
Glu Phe Xaa Gly Pro Ser Gly Leu Leu Thr Tyr Thr Gly Lys Gly His
          120          125          130
Phe Asn Ile Gln Pro Asn Lys Asn Leu His Gln Asn Pro Glu Trp Arg
135          140          145          150
Arg Asn Trp Glu

```

<210> 455

<211> 91

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -64...-1

<400> 455

```

Met Thr Pro Arg Ile Leu Ser Glu Val Gln Phe Ser Ala Phe Cys Pro
          -60          -55          -50
Tyr Trp Thr Ile Ala Arg Ile Leu Glu Arg Val Gly Ser Ala Cys Phe
          -45          -40          -35
Arg Leu Glu Leu Cys Ala Ala Ile Val Gly Tyr Phe Val Leu Asp Val
          -30          -25          -20
Arg Thr Phe Leu Phe Ile Val Val Cys Val Ile Cys Val Thr Leu Asn
          -15          -10          -5
Phe Pro Arg Phe Tyr Phe Leu Cys Leu Ser Ser Leu Thr Ala Phe Gly
1          5          10          15
Thr Pro Pro Ile Gly Val His Ile Pro Ser Pro
          20          25

```

<210> 456

<211> 257

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -23...-1

<400> 456

Met	Arg	Arg	Ile	Ser	Leu	Thr	Ser	Ser	Pro	Val	Arg	Leu	Leu	Leu	Xaa
			-20					-15					-10		
Leu	Leu	Leu	Leu	Leu	Ile	Ala	Leu	Glu	Ile	Met	Val	Gly	Gly	His	Ser
		-5					1				5				
Leu	Cys	Phe	Asn	Phe	Thr	Ile	Lys	Ser	Leu	Ser	Arg	Pro	Gly	Gln	Pro
10					15				20					25	
Trp	Cys	Glu	Ala	His	Val	Phe	Leu	Asn	Lys	Asn	Leu	Phe	Leu	Gln	Tyr
				30				35						40	
Asn	Ser	Asp	Asn	Asn	Met	Val	Lys	Pro	Leu	Gly	Leu	Leu	Gly	Lys	Lys
			45				50						55		
Val	Tyr	Ala	Thr	Ser	Thr	Trp	Gly	Glu	Leu	Thr	Gln	Thr	Leu	Gly	Glu
		60					65				70				
Val	Gly	Arg	Asp	Leu	Arg	Met	Leu	Leu	Cys	Asp	Ile	Lys	Pro	Gln	Ile
		75				80				85					
Lys	Thr	Ser	Asp	Pro	Ser	Thr	Leu	Gln	Val	Xaa	Xaa	Phe	Cys	Gln	Arg
90					95				100						105
Glu	Ala	Glu	Arg	Cys	Thr	Gly	Ala	Ser	Trp	Gln	Phe	Ala	Thr	Asn	Gly
				110					115					120	
Glu	Lys	Ser	Leu	Leu	Phe	Asp	Ala	Met	Asn	Met	Thr	Trp	Thr	Val	Ile
			125				130						135		
Asn	His	Glu	Ala	Ser	Xaa	Ile	Lys	Glu	Thr	Trp	Lys	Lys	Asp	Arg	Xaa
			140				145					150			
Leu	Glu	Xaa	Tyr	Phe	Arg	Lys	Leu	Ser	Lys	Gly	Asp	Cys	Asp	His	Trp
		155				160				165					
Leu	Arg	Glu	Phe	Leu	Gly	His	Trp	Glu	Ala	Met	Pro	Xaa	Pro	Xaa	Val
170					175				180						185
Ser	Pro	Xaa	Asn	Ala	Ser	Xaa	Ile	His	Trp	Ser	Ser	Ser	Xaa	Leu	Pro
			190					195						200	
Xaa	Xaa	Trp	Ile	Ile	Leu	Gly	Ala	Phe	Ile	Leu	Leu	Xaa	Leu	Met	Gly
			205				210						215		
Ile	Val	Leu	Ile	Cys	Val	Trp	Trp	Gln	Asn	Gly	Xaa	Xaa	Ser	Thr	Xaa
		220					225						230		
Xaa															

<210> 457

<211> 193

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -60..-1

<400> 457

Met	Cys	Pro	Ser	Leu	Glu	Glu	Ala	Pro	Ser	Val	Lys	Gly	Thr	Leu	Pro
-60					-55					-50				-45	
Cys	Ser	Gly	Gln	Gln	Gln	Pro	Phe	Pro	Phe	Gly	Ala	Ser	Asn	Ile	Pro
			-40					-35						-30	
Leu	Leu	Leu	Gly	Arg	Ser	Arg	Lys	Val	Ala	Arg	Gly	Ala	Pro	Val	Leu
		-25					-20						-15		
Trp	Pro	Phe	Leu	Thr	Trp	Ile	Asn	Pro	Ala	Leu	Ser	Ile	Cys	Asp	Pro
	-10					-5					1				
Leu	Gly	Ser	Cys	Gly	Trp	Xaa	Cys	His	Thr	Ala	Gln	Val	Pro	Ala	Pro

```

5          10          15          20
Leu Gln Leu Pro Thr Ala Cys Pro Pro Leu Pro His Gly Thr Arg Ala
          25          30          35
Val Gly Pro Thr Pro Gly Leu Leu Pro Glu Ala Ala Ala Pro Xaa Thr
          40          45          50
Xaa Gly Ala Leu Ser Ser Arg Ser Arg His Trp Ser Cys Ser Ile Val
          55          60          65
Xaa Cys Leu His Leu His Xaa Leu Leu Ser Val Glu Thr Arg Xaa Phe
          70          75          80
Xaa Lys His Leu Leu Val Leu Leu Val Ala Val Ala His Ser Val Leu
85          90          95          100
Glu Pro Pro Ala Leu Val Pro Asn Val Gln Cys Glu Met Cys Thr His
          105          110          115
Ser Gly Pro Arg Asp Leu Glu Ala Ala Val Val Ser Pro Ala Pro Trp
          120          125          130
Glu

```

```

<210> 458
<211> 107
<212> PRT
<213> Homo sapiens

```

```

<220>
<221> SIGNAL
<222> -28...-1

```

```

<400> 458
Met Val Leu Thr Leu Gly Glu Ser Trp Pro Val Leu Val Gly Arg Arg
          -25          -20          -15
Phe Leu Ser Leu Ser Ala Ala Asp Gly Ser Asp Gly Ser His Asp Ser
          -10          -5          1
Trp Asp Val Glu Arg Val Ala Glu Trp Pro Trp Leu Ser Gly Thr Ile
5          10          15          20
Arg Ala Val Ser His Thr Asp Val Thr Lys Lys Asp Leu Lys Val Cys
          25          30          35
Val Glu Phe Xaa Gly Glu Ser Trp Arg Lys Arg Arg Trp Ile Glu Val
          40          45          50
Tyr Ser Leu Leu Arg Lys Ala Phe Leu Val Lys His Asn Leu Val Leu
          55          60          65
Ala Glu Arg Lys Ser Pro Glu Ile Ser Trp Gly
70          75

```

```

<210> 459
<211> 121
<212> PRT
<213> Homo sapiens

```

```

<220>
<221> SIGNAL
<222> -13...-1

```

```

<400> 459
Met Leu Val Leu Arg Ser Ala Leu Thr Arg Ala Leu Ala Ser Arg Thr

```

```

      -10      -5      1
Leu Ala Pro Gln Met Cys Ser Ser Phe Ala Thr Gly Pro Arg Gln Tyr
  5          10      15
Asp Gly Ile Phe Tyr Glu Phe Arg Ser Tyr Tyr Leu Lys Pro Ser Lys
20          25      30      35
Met Asn Glu Phe Leu Glu Asn Phe Glu Lys Asn Ala Gln Leu Arg Thr
      40          45      50
Ala His Ser Glu Leu Val Gly Tyr Trp Ser Val Xaa Phe Gly Gly Arg
      55          60      65
Met Xaa Thr Val Phe His Ile Trp Lys Tyr Asp Asn Phe Ala His Arg
      70          75      80
Thr Glu Phe Gln Lys Ala Leu Ala Lys Asp Lys Glu Trp Gln Glu Gln
      85          90      95
Phe Leu Ile Pro Asn Leu Ala Leu Asn
100          105

```

<210> 460

<211> 44

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -17...-1

<400> 460

```

Met Lys Val Gly Val Leu Trp Leu Ile Ser Phe Phe Thr Phe Thr Asp
      -15      -10      -5
Gly His Gly Gly Phe Leu Gly Val Ser Trp Cys Tyr Val Ser Tyr Leu
  1          5      10      15
Phe Ser Thr Asn Ser Pro Leu Ser Phe Arg Arg Ile
      20          25

```

<210> 461

<211> 109

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -13...-1

<400> 461

```

Met Cys Leu Leu Thr Ala Leu Val Thr Gln Val Ile Ser Leu Arg Lys
      -10      -5      1
Asn Ala Glu Arg Thr Cys Leu Cys Lys Arg Arg Trp Pro Trp Xaa Pro
  5          10      15
Ser Pro Arg Ile Tyr Cys Ser Ser Thr Pro Cys Asp Ser Lys Phe Pro
20          25      30      35
Thr Val Tyr Ser Ser Ala Pro Phe His Ala Pro Leu Pro Val Gln Asn
      40          45      50
Ser Leu Trp Gly His Pro Leu His Gly Cys Ser Trp Gln Cys His His
      55          60      65

```



Pro	Gln	Gly	Gln	Asn	Leu	Gln	Pro	Ala	Ser	Leu	Xaa	Thr	His	Leu	Ser
	70						75					80			
Lys	Pro	Lys	Arg	His	Phe	Xaa	Lys	Lys	Xaa	Cys	Gln	Ala			
85						90					95				

<210> 462  
 <211> 143  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -41..-1

Met	Ala	Thr	Ala	Thr	Glu	Gln	Trp	Val	Leu	Val	Glu	Met	Val	Gln	Ala
-40						-35					-30				
Leu	Tyr	Glu	Ala	Pro	Ala	Tyr	His	Leu	Ile	Leu	Glu	Gly	Ile	Leu	Ile
-25					-20					-15					-10
Leu	Trp	Ile	Ile	Arg	Leu	Leu	Phe	Ser	Lys	Thr	Tyr	Lys	Leu	Gln	Glu
				-5					1				5		
Arg	Ser	Asp	Leu	Thr	Val	Lys	Glu	Lys	Glu	Glu	Leu	Ile	Glu	Glu	Trp
		10				15						20			
Gln	Pro	Glu	Pro	Leu	Val	Pro	Pro	Val	Pro	Lys	Asp	His	Pro	Ala	Leu
	25					30					35				
Asn	Tyr	Asn	Ile	Val	Ser	Gly	Pro	Pro	Ser	His	Lys	Thr	Val	Val	Asn
40					45					50					55
Gly	Lys	Glu	Cys	Ile	Asn	Phe	Ala	Ser	Phe	Asn	Phe	Leu	Gly	Leu	Leu
				60					65					70	
Asp	Asn	Pro	Arg	Val	Lys	Ala	Ala	Ala	Leu	Ala	Ser	Leu	Lys	Lys	Tyr
			75					80					85		
Gly	Val	Gly	Thr	Cys	Gly	Pro	Cys	Gly	Phe	Tyr	Gly	Thr	Phe	Glu	
	90						95					100			

<210> 463  
 <211> 232  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -30..-1

Met	Ala	Ala	Thr	Ser	Gly	Thr	Asp	Glu	Pro	Val	Ser	Gly	Glu	Leu	Val
-30					-25					-20					-15
Ser	Val	Ala	His	Ala	Leu	Ser	Leu	Pro	Ala	Glu	Ser	Tyr	Gly	Asn	Xaa
			-10						-5					1	
Xaa	Asp	Ile	Glu	Met	Ala	Trp	Ala	Met	Arg	Ala	Met	Gln	His	Ala	Glu
	5					10						15			
Val	Tyr	Tyr	Lys	Leu	Ile	Ser	Ser	Val	Asp	Pro	Gln	Phe	Leu	Lys	Leu
	20					25					30				
Thr	Lys	Val	Asp	Asp	Gln	Ile	Tyr	Ser	Glu	Phe	Arg	Lys	Asn	Phe	Glu

35		40		45		50									
Thr	Leu	Arg	Ile	Asp	Val	Leu	Xaa	Pro	Glu	Xaa	Leu	Lys	Ser	Glu	Ser
				55					60					65	
Ala	Lys	Glu	Pro	Pro	Gly	Tyr	Asn	Ser	Leu	Pro	Leu	Lys	Leu	Leu	Gly
			70					75					80		
Thr	Gly	Lys	Ala	Ile	Thr	Lys	Leu	Phe	Ile	Ser	Val	Phe	Arg	Thr	Lys
		85					90					95			
Lys	Glu	Arg	Lys	Glu	Ser	Thr	Met	Glu	Glu	Lys	Lys	Glu	Leu	Thr	Val
	100					105					110				
Glu	Lys	Lys	Arg	Thr	Pro	Arg	Met	Glu	Glu	Arg	Lys	Glu	Leu	Ile	Val
	115				120					125				130	
Glu	Lys	Lys	Lys	Arg	Lys	Glu	Ser	Thr	Glu	Lys	Thr	Lys	Leu	Thr	Lys
			135						140				145		
Glu	Glu	Lys	Lys	Gly	Lys	Lys	Leu	Thr	Lys	Lys	Ser	Thr	Lys	Val	Val
		150					155					160			
Lys	Lys	Leu	Cys	Lys	Val	Tyr	Arg	Glu	Gln	His	Ser	Arg	Ser	Tyr	Asp
	165					170					175				
Ser	Ile	Glu	Thr	Thr	Ser	Thr	Thr	Val	Leu	Leu	Ala	Gln	Thr	Pro	Leu
	180					185					190				
Val	Lys	Cys	Lys	Phe	Leu	Tyr	Asn								
	195				200										

<210> 464

<211> 61

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -21...-1

<400> 464

Met	Thr	Phe	Arg	His	Gln	Asp	Asn	Ser	Leu	Met	Phe	Phe	Ser	Met	Met
	-20					-15					-10				
Ala	Thr	Cys	Thr	Ser	Asn	Val	Gly	Phe	Thr	His	Thr	Thr	Met	Asn	Cys
	5				1				5					10	
Ser	Leu	Thr	Ser	Pro	Val	Asp	Phe	Lys	Asp	Leu	Leu	Arg	Val	Leu	Leu
		15				20							25		
Ile	Lys	Phe	Gly	Tyr	Asp	Arg	Lys	Ser	Thr	Ile	Lys	Ser			
	30					35						40			

<210> 465

<211> 34

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -19...-1

<400> 465

Met	Phe	Leu	Lys	Ser	Gly	Ala	Gly	Leu	Ser	Ser	Cys	Leu	Leu	Pro	Leu
				-15					-10					-5	

Cys Trp Leu Glu Arg Lys Asp His Gly Arg Arg Pro Ser Xaa His Pro  
1 5 10  
Gly Arg  
15

<210> 466  
<211> 215  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -54..-1

<400> 466  
Met Asn Xaa Tyr Ala Ser Pro Phe Asn Xaa Gln Leu Xaa Tyr Leu Xaa  
-50 -45 -40  
Leu Ser Arg Phe Glu Cys Val His Arg Asp Gly Arg Val Ile Thr Leu  
-35 -30 -25  
Ser Tyr Gln Glu Gln Glu Leu Gln Asp Phe Leu Leu Ser Gln Met Ser  
-20 -15 -10  
Gln His Gln Val His Ala Val Gln Gln Leu Ala Lys Val Met Gly Trp  
-5 1 5 10  
Gln Val Leu Ser Phe Ser Asn His Val Gly Leu Gly Pro Ile Glu Ser  
15 20 25  
Xaa Gly Asn Ala Ser Ala Ile Thr Val Ala Pro Gln Val Val Thr Met  
30 35 40  
Leu Phe Gln Phe Val Met Asp Leu Lys Val Ala Ala Arg Leu Trp Phe  
45 50 55  
Ser Phe Leu Val Thr Asn Val Lys Thr Phe Gln Lys Val Met Phe Tyr  
60 65 70  
Lys Ile Thr Asn Gly Val Ile Phe Val Gly His Ser Lys Lys Phe Ser  
75 80 85 90  
Gly Ile Lys Trp Lys Val Xaa Ile Leu Phe Ile Lys Trp Xaa Cys Leu  
95 100 105  
Cys Leu His Leu Ala Leu Val Tyr Tyr Asp Phe Phe Gln Met Phe Pro  
110 115 120  
Lys Xaa Val Ser Xaa Asn Phe Asp Leu Lys Cys Leu Gln Ile Asn Tyr  
125 130 135  
Lys His Lys Glu Glu Ile Thr Ser Lys Arg Val Leu Phe Leu Lys Ile  
140 145 150  
Ile Ile Arg Lys Cys Phe Ile  
155 160

<210> 467  
<211> 27  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -17..-1

<400> 467

Met Val Val His Leu Leu Tyr Ala His Leu Ser Phe Thr Ser Lys Arg  
 -15 -10 -5  
 Ala Val Val Met Leu Lys Leu Glu Ile Thr Phe  
 1 5 10

<210> 468

<211> 85

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -24..-1

<400> 468

Met Cys Ser His Ala Ser Met Ser Phe His Thr Leu Phe His Leu Leu  
 -20 -15 -10  
 Phe Leu Pro His Tyr Ile Glu Thr Phe Lys Pro Gln Ser Lys His Cys  
 -5 1 5  
 Phe Phe Trp Ile Ala Ala Phe Leu Thr Ser Leu Leu Thr Pro Gln Ser  
 10 15 20  
 Leu Gln Gly Phe His Ser Ser Leu Cys Ala Leu Arg Ser Gln His Phe  
 25 30 35 40  
 Pro Ser Thr Cys Asn Cys Phe Cys Tyr Leu Thr Ile Ile Ala Leu Xaa  
 45 50 55  
 Tyr Trp Asp Asn Leu  
 60

<210> 469

<211> 51

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -16..-1

<400> 469

Met Leu Arg Ile Ala Leu Thr Leu Ile Pro Ser Met Leu Ser Arg Ala  
 -15 -10 -5  
 Ala Gly Trp Cys Trp Tyr Lys Glu Pro Thr Gln Gln Phe Ser Tyr Leu  
 1 5 10 15  
 Cys Leu Pro Cys Leu Ser Trp Asn Lys Lys Gly Asn Val Leu Gln Leu  
 20 25 30  
 Pro Asn Phe  
 35

<210> 470

<211> 67

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -43..-1

<400> 470

```

Met Thr Pro Gln Tyr Leu Pro His Gly Gly Lys Tyr Gln Val Leu Gly
      -40                      -35                      -30
Asp Tyr Ser Leu Ala Val Val Phe Pro Leu His Phe Ser Asp Leu Ile
      -25                      -20                      -15
Ser Val Leu Tyr Leu Ile Pro Lys Thr Leu Thr Thr Asn Thr Ala Val
      -10                      -5                      1                      5
Lys His Ser Ile Gln Lys Asn Cys Met Xaa Leu Val Leu Gly Lys Leu
      10                      15                      20
Leu Ser Gln

```

<210> 471

<211> 63

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -15..-1

<400> 471

```

Met Gly Ile Leu Ser Thr Val Thr Ala Leu Thr Phe Ala Arg Ala Leu
      -15                      -10                      -5                      1
Asp Gly Cys Arg Asn Gly Ile Ala His Pro Ala Ser Glu Lys His Arg
      5                      10                      15
Leu Glu Lys Cys Arg Glu Leu Glu Ser Ser His Ser Ala Pro Gly Ser
      20                      25                      30
Thr Gln His Arg Arg Lys Thr Thr Arg Arg Asn Tyr Ser Ser Ala
      35                      40                      45

```

<210> 472

<211> 179

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -58..-1

<400> 472

```

Met Ser Thr Gly Gln Leu Tyr Arg Met Glu Asp Ile Gly Arg Phe His
      -55                      -50                      -45
Ser Gln Gln Pro Gly Ser Leu Thr Pro Ser Ser Pro Thr Val Gly Glu
      -40                      -35                      -30
Ile Ile Tyr Asn Asn Thr Arg Asn Thr Leu Gly Trp Ile Gly Gly Ile
      -25                      -20                      -15
Leu Met Gly Ser Phe Gln Gly Thr Ile Ala Gly Gln Gly Thr Gly Ala

```

-10		-5		1		5											
Thr	Ser	Ile	Ser	Glu	Leu	Cys	Lys	Gly	Gln	Glu	Leu	Glu	Pro	Ser	Gly		
			10					15					20				
Ala	Gly	Leu	Thr	Val	Ala	Pro	Pro	Gln	Ala	Val	Ser	Leu	Gln	Gly	Ile		
		25					30					35					
Tyr	Thr	Leu	Pro	Trp	Leu	Leu	Gln	Leu	Phe	His	Ser	Thr	Ala	Leu	Xaa		
		40				45					50						
Xaa	Xaa	Gln	Gln	Pro	Asn	Gly	Ser	Leu	Ser	Leu	Asn	Ile	Ser	Ser	Ser		
55					60					65				70			
His	Ala	Pro	Xaa	Pro	Xaa	Thr	Cys	Thr	Leu	Glu	Pro	Gly	Val	Asp	Pro		
				75					80					85			
Thr	Arg	Xaa	Val	Cys	Ile	Asn	Pro	His	Pro	Pro	Pro	Pro	Ile	Leu	Lys		
			90					95					100				
Xaa	Pro	Leu	Ser	Pro	Tyr	Pro	Lys	Pro	Gln	Leu	Gly	Thr	His	Ala	Gly		
		105					110					115					
Gln	Val	Asn															
		120															

<210> 473  
 <211> 238  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -71...-1

<400> 473																	
Met	Xaa	Xaa	Phe	Thr	Asp	Pro	Ser	Ser	Val	Asn	Glu	Lys	Lys	Arg	Arg		
	-70				-65					-60							
Glu	Arg	Glu	Glu	Arg	Gln	Asn	Ile	Val	Leu	Trp	Arg	Gln	Pro	Leu	Ile		
55					-50					-45				-40			
Thr	Leu	Gln	Tyr	Phe	Ser	Leu	Glu	Ile	Leu	Val	Ile	Leu	Lys	Glu	Trp		
				-35					-30					-25			
Thr	Ser	Lys	Leu	Trp	His	Arg	Gln	Ser	Ile	Val	Val	Ser	Phe	Leu	Leu		
			-20				-15						-10				
Leu	Leu	Ala	Gly	Leu	Ile	Ala	Thr	Tyr	Tyr	Val	Glu	Gly	Val	His	Gln		
	-5					1			5								
Gln	Tyr	Val	Gln	Arg	Ile	Glu	Lys	Gln	Phe	Leu	Leu	Tyr	Ala	Tyr	Trp		
10				15					20				25				
Ile	Gly	Leu	Gly	Ile	Leu	Ser	Ser	Val	Gly	Leu	Gly	Thr	Gly	Leu	His		
			30					35					40				
Thr	Phe	Leu	Leu	Tyr	Leu	Gly	Pro	His	Ile	Ala	Ser	Val	Thr	Leu	Ala		
		45					50					55					
Ala	Tyr	Glu	Cys	Asn	Ser	Val	Asn	Phe	Pro	Glu	Pro	Pro	Tyr	Pro	Asp		
	60					65						70					
Gln	Ile	Ile	Cys	Pro	Asp	Glu	Gly	Thr	Glu	Gly	Thr	Ile	Ser	Leu			
	75				80					85							
Trp	Ser	Ile	Ile	Ser	Lys	Val	Arg	Ile	Glu	Ala	Cys	Met	Trp	Gly	Ile		
90					95				100					105			
Gly	Thr	Ala	Ile	Gly	Glu	Leu	Pro	Pro	Tyr	Phe	Met	Ala	Arg	Ala	Ala		
			110					115					120				
Arg	Leu	Ser	Gly	Ala	Glu	Pro	Asp	Asp	Glu	Glu	Tyr	Gln	Glu	Phe	Glu		
		125					130					135					
Glu	Met	Leu	Glu	His	Ala	Glu	Ser	Ala	Gln	Val	Arg	Thr	Val	Gly	Ile		

	140						145					150	
Glu	Asn	Arg	Thr	Leu	Tyr	Phe	Phe	Leu	Lys	Arg	Leu	Leu	Arg
155						160					165		

<210> 474  
 <211> 178  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -37..-1

<400> 474

Met	Glu	Arg	Gln	Ser	Arg	Val	Met	Ser	Glu	Lys	Asp	Glu	Tyr	Gln	Phe
	-35						-30					-25			
Gln	His	Gln	Gly	Ala	Val	Glu	Leu	Leu	Val	Phe	Asn	Phe	Leu	Leu	Ile
-20						-15					-10				
Leu	Thr	Ile	Leu	Thr	Ile	Trp	Leu	Phe	Lys	Asn	His	Arg	Phe	Arg	Phe
5					1				5					10	
Leu	His	Glu	Thr	Gly	Gly	Ala	Met	Val	Tyr	Gly	Leu	Xaa	Met	Gly	Leu
		15					20					25			
Ile	Leu	Xaa	Tyr	Ala	Thr	Ala	Pro	Thr	Asp	Ile	Glu	Ser	Gly	Xaa	Val
	30					35					40				
Tyr	Asp	Cys	Val	Lys	Leu	Thr	Phe	Ser	Pro	Ser	Thr	Leu	Leu	Val	Asn
45					50					55					
Ile	Thr	Asp	Gln	Val	Tyr	Glu	Tyr	Lys	Tyr	Lys	Arg	Glu	Ile	Ser	Gln
60					65					70					75
His	Xaa	Ile	Asn	Pro	His	Xaa	Gly	Asn	Ala	Ile	Leu	Glu	Lys	Met	Thr
		80					85							90	
Phe	Asp	Pro	Xaa	Ile	Phe	Phe	Asn	Val	Leu	Leu	Pro	Pro	Ile	Ile	Phe
		95					100					105			
His	Ala	Gly	Tyr	Ser	Leu	Lys	Lys	Arg	His	Phe	Phe	Gln	Asn	Leu	Gly
	110					115						120			
Ser	Ile	Leu	Thr	Tyr	Ala	Phe	Leu	Gly	Thr	Ala	Ile	Ser	Cys	Ile	Val
125						130					135				
Ile	Gly														
140															

<210> 475  
 <211> 96  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -21..-1

<400> 475

Met	Ser	Met	Gln	Phe	Leu	Phe	Lys	Met	Val	Ala	Leu	Cys	Cys	Cys	Leu
-20						-15					-10				
Trp	Lys	Ile	Ser	Gly	Cys	Glu	Glu	Val	Pro	Leu	Thr	Tyr	Asn	Leu	Leu
-5				1				5						10	

Lys	Cys	Leu	Leu	Asp	Lys	Ala	His	Cys	Val	Leu	Leu	Thr	Pro	Cys	Gly
			15					20					25		
Tyr	Ile	Phe	Ser	Leu	Ile	Ser	Pro	Glu	Ile	Leu	Lys	Leu	Thr	Leu	Ile
		30					35					40			
Thr	Leu	Xaa	Ile	Leu	Leu	Ile	Leu	Lys	Asn	Leu	His	Leu	Leu	Trp	Leu
	45					50					55				
Thr	Val	Ser	Ser	Xaa	Cys	Val	His	Arg	Ser	Ser	Ala	Arg	Lys	Glu	Lys
60					65					70					75

```
<210> 476
<211> 41
<212> PRT
<213> Homo sapiens
```

```
<220>
<221> SIGNAL
<222> -24...-1
```

```

<400> 476
Met His Thr Phe Ala Asn Asp Arg Gly Leu Tyr Arg Ile Leu Leu Leu
          -20          -15          -10
His Phe Tyr Cys Leu Leu Arg Ser Ser Glu Tyr Ile Leu Gly Tyr Lys
          -5          1          5
Val Leu Gly Val Phe Phe Pro Ile Leu
    10          15

```

```
<210> 477
<211> 113
<212> PRT
<213> Homo sapiens
```

```

<220>
<221> SIGNAL
<222> -27..-1

```

<400>																			
Met	Arg	Xaa	Lys	Trp	Lys	Met	Gly	Gly	Met	Lys	Tyr	Ile	Phe	Ser	Leu				
-25								-20									-15		
Leu	Phe	Phe	Leu	Leu	Leu	Glu	Gly	Gly	Xaa	Thr	Glu	Gln	Val	Xaa	His				
-10								-5									1	5	
Ser	Glu	Thr	Tyr	Cys	Met	Phe	Gln	Asp	Lys	Lys	Tyr	Arg	Val	Gly	Glu				
							10								15			20	
Arg	Trp	His	Pro	Tyr	Leu	Glu	Pro	Tyr	Gly	Leu	Val	Tyr	Cys	Val	Asn				
							25								30			35	
Cys	Ile	Cys	Ser	Glu	Asn	Gly	Asn	Val	Leu	Cys	Ser	Arg	Val	Arg	Cys				
							40								45			50	
Pro	Asn	Val	His	Cys	Leu	Ser	Pro	Val	His	Ile	Pro	His	Leu	Cys	Cys				
							55								60			65	
Pro	Arg	Cys	Pro	Glu	Asp	Ser	Leu	Pro	Pro	Val	Asn	Asn	Xaa	Val	Thr				
70								75								80			85
Ser																			



<210> 478  
 <211> 250  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -18..-1

<400> 478  
 Met Arg Ile Leu Gln Leu Ile Leu Leu Ala Leu Ala Thr Gly Leu Val  
                   -15                  -10                  -5  
 Gly Gly Glu Thr Arg Ile Ile Lys Gly Phe Glu Cys Lys Pro His Ser  
           1                  5                  10  
 Gln Pro Trp Gln Ala Ala Leu Phe Glu Lys Thr Arg Leu Leu Cys Gly  
 15                  20                  25                  30  
 Ala Thr Leu Ile Ala Pro Arg Trp Leu Leu Thr Ala Ala His Cys Leu  
                   35                  40                  45  
 Lys Pro Arg Tyr Ile Xaa His Leu Gly Gln His Asn Leu Gln Lys Glu  
                   50                  55                  60  
 Glu Gly Cys Glu Gln Thr Arg Thr Ala Thr Glu Ser Phe Pro His Pro  
           65                  70                  75  
 Gly Phe Asn Asn Ser Leu Pro Asn Lys Asp Xaa Xaa Asn Asp Ile Met  
           80                  85                  90  
 Leu Val Xaa Met Xaa Ser Pro Val Ser Ile Thr Trp Ala Val Arg Pro  
 95                  100                  105                  110  
 Leu Thr Leu Ser Ser Arg Cys Val Thr Ala Gly Thr Ser Cys Leu Ile  
                   115                  120                  125  
 Ser Gly Trp Gly Ser Thr Ser Ser Pro Gln Leu Arg Leu Pro His Thr  
           130                  135                  140  
 Leu Arg Cys Ala Asn Ile Thr Ile Ile Glu His Gln Lys Cys Glu Asn  
           145                  150                  155  
 Ala Tyr Pro Gly Asn Ile Thr Asp Thr Met Val Cys Ala Ser Val Gln  
           160                  165                  170  
 Glu Gly Gly Lys Asp Ser Cys Gln Gly Asp Ser Gly Gly Pro Leu Val  
 175                  180                  185                  190  
 Cys Asn Gln Ser Leu Gln Gly Ile Ile Ser Trp Gly Gln Asp Pro Cys  
                   195                  200                  205  
 Ala Ile Thr Arg Lys Pro Gly Val Tyr Thr Lys Val Cys Lys Tyr Val  
           210                  215                  220  
 Asp Trp Ile Gln Glu Thr Met Lys Asn Asn  
           225                  230

<210> 479  
 <211> 151  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -21..-1

<400> 479  
 Met Ala Ala Ser Thr Ser Met Val Pro Val Ala Val Thr Ala Ala Val

-20		-15		-10
Ala Pro Val Leu Ser Ile Asn Ser Asp Phe Ser Asp Leu Arg Glu Ile				
-5		1	5	10
Lys Lys Gln Leu Leu Leu Ile Ala Gly Leu Thr Arg Glu Arg Gly Leu				
	15		20	25
Leu His Ser Ser Lys Trp Ser Ala Glu Leu Ala Phe Ser Leu Pro Ala				
	30	35	40	
Leu Pro Leu Ala Glu Leu Gln Pro Pro Pro Pro Ile Thr Glu Glu Asp				
	45	50	55	
Ala Gln Asp Met Asp Ala Tyr Thr Leu Ala Lys Ala Tyr Phe Asp Val				
60		65	70	75
Lys Glu Tyr Asp Arg Ala Ala His Phe Leu His Gly Cys Asn Ala Arg				
	80	85	90	
Lys Ala Tyr Phe Leu Tyr Met Tyr Ser Arg Tyr Leu Val Arg Ala Ile				
	95	100	105	
Leu Lys Cys His Ser Ala Phe Ser Glu Thr Ser Ile Phe Arg Thr Asn				
	110	115	120	
Gly Lys Val Lys Ser Phe Lys				
	125	130		

A210> 480  
 A211> 239  
 A212> PRT  
 A213> Homo sapiens

A220>  
 A221> SIGNAL  
 A222> -25...-1

A400> 480  
 Met Pro Arg Lys Arg Lys Cys Asp Leu Arg Ala Val Arg Val Gly Leu  
 -25 -20 -15 -10  
 Leu Leu Gly Gly Gly Gly Val Tyr Gly Ser Arg Phe Arg Phe Thr Phe  
 -5 1 5  
 Pro Gly Cys Arg Ala Leu Ser Pro Trp Arg Val Arg Xaa Gln Arg Arg  
 10 15 20  
 Arg Cys Glu Met Ser Thr Met Phe Ala Asp Thr Leu Leu Ile Val Phe  
 25 30 35  
 Ile Ser Val Cys Thr Ala Leu Leu Ala Glu Gly Ile Thr Trp Val Leu  
 40 45 50 55  
 Val Tyr Arg Thr Asp Lys Tyr Lys Arg Leu Lys Ala Glu Val Glu Lys  
 60 65 70  
 Gln Ser Lys Lys Leu Glu Lys Lys Lys Glu Thr Ile Thr Glu Ser Ala  
 75 80 85  
 Gly Arg Gln Gln Lys Lys Lys Ile Glu Arg Xaa Xaa Xaa Xaa Leu Xaa  
 90 95 100  
 Asn Asn Asn Arg Asp Leu Ser Met Val Arg Met Lys Ser Met Phe Ala  
 105 110 115  
 Ile Gly Phe Cys Phe Thr Ala Leu Met Gly Met Phe Asn Ser Ile Phe  
 120 125 130 135  
 Asp Gly Arg Val Val Ala Lys Leu Pro Phe Thr Pro Leu Ser Xaa Xaa  
 140 145 150  
 Xaa Gly Leu Ser His Arg Asn Leu Leu Gly Asp Asp Thr Thr Asp Cys  
 155 160 165  
 Ser Phe Ile Phe Leu Xaa Ile Leu Cys Thr Met Ser Ile Arg Gln Asn

-371-

170	175	180
Ile Gln Lys Ile Leu Gly	Leu Ala Pro Ser Arg	Ala Ala Thr Lys Gln
185	190	195
Ala Gly Gly Phe Leu Gly	Pro Pro Pro Pro Ser	Gly Lys Phe Ser
200	205	210

<210> 481  
<211> 208  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -92..-1

<400> 481	
Met Arg Glu Pro Gln Lys Arg Thr Ala Thr Ile Ala Lys Xaa Xaa Ala	
-90	-85 -80
Xaa Glu Gly Leu Arg Asp Pro Tyr Gly Arg Leu Cys Gly Ser Glu His	
-75	-70 -65
Pro Arg Arg Pro Pro Glu Arg Pro Glu Glu Asp Pro Ser Thr Pro Glu	
60	-55 -50 -45
Glu Ala Ser Thr Thr Pro Glu Glu Ala Ser Ser Thr Ala Gln Ala Gln	
	-40 -35 -30
Lys Pro Ser Val Pro Arg Ser Asn Phe Gln Gly Thr Lys Lys Ser Leu	
-25	-20 -15
Leu Met Ser Ile Leu Ala Leu Ile Phe Ile Met Gly Asn Ser Ala Lys	
-10	-5 1
Glu Ala Leu Val Trp Lys Val Leu Gly Lys Leu Gly Met Gln Pro Gly	
5	10 15 20
Arg Xaa His Ser Ile Phe Gly Asp Pro Lys Lys Ile Val Thr Glu Xaa	
25	30 35
Phe Val Arg Arg Gly Tyr Leu Ile Tyr Xaa Pro Val Pro Arg Xaa Ser	
40	45 50
Pro Val Glu Tyr Xaa Phe Phe Trp Gly Pro Arg Ala His Val Glu Ser	
55	60 65
Ser Xaa Leu Lys Xaa Xaa His Phe Val Ala Arg Val Arg Asn Arg Cys	
70	75 80
Ser Lys Asp Trp Pro Cys Asn Tyr Asp Trp Asp Ser Asp Asp Ala	
85	90 95 100
Glu Val Glu Ala Ile Leu Asn Ser Gly Ala Xaa Gly Tyr Ser Ala Pro	
	105 110 115

<210> 482  
<211> 86  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -39..-1

<400> 482

```

Met Asn Val Gly Thr Ala His Xaa Xaa Val Asn Pro Asn Thr Arg Val
      -35                      -30                      -25
Met Asn Ser Arg Gly Ile Trp Leu Ser Tyr Val Leu Ala Ile Gly Leu
      -20                      -15                      -10
Leu His Ile Val Leu Leu Ser Ile Pro Phe Val Ser Val Pro Val Val
      -5                      1                      5
Trp Thr Leu Thr Asn Leu Ile His Asn Met Gly Met Tyr Ile Phe Leu
10      15      20      25
His Thr Val Lys Gly Thr Pro Phe Glu Thr Pro Asp Gln Gly Lys Ala
      30                      35                      40
Arg Leu Leu Thr His Trp
      45

```

<210> 483  
 <211> 40  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -27...-1

```

<400> 483
Met Arg Thr Leu Phe Gly Ala Val Arg Ala Pro Phe Ser Ser Leu Thr
      -25                      -20                      -15
Leu Leu Leu Ile Thr Pro Ser Pro Ser Pro Leu Leu Phe Asp Arg Gly
      -10                      -5                      1                      5
Leu Ser Leu Arg Ser Ala Met Ser
      10

```

<210> 484  
 <211> 65  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -16...-1

```

<400> 484
Met Leu Gly Phe Phe Leu Phe Leu Ser Phe Val Leu Met Tyr Asp Gly
      -15                      -10                      -5
Leu Arg Leu Phe Gly Ile Leu Ser Thr Cys Arg Val His His Thr Met
1      5      10      15
Asn Gln Phe Leu Ile Asp Ile Ser Ser Phe Thr Ser Arg Val Lys Lys
      20      25      30
Lys Ile Phe Leu Phe Tyr Ala Phe Xaa Gly Cys Xaa Phe Gln Ser Ala
      35      40      45
Thr

```

<210> 485

<211> 130  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -55..-1

<400> 485  
Met Ala Met Trp Asn Arg Pro Xaa Xaa Xaa Leu Pro Gln Gln Pro Leu  
-55 -50 -45 -40  
Xaa Ala Glu Pro Thr Ala Glu Gly Glu Pro His Leu Pro Thr Gly Arg  
-35 -30 -25  
Xaa Xaa Thr Glu Ala Asn Arg Phe Ala Tyr Ala Ala Leu Cys Gly Ile  
-20 -15 -10  
Ser Leu Ser Gln Leu Phe Pro Glu Pro Glu His Ser Ser Phe Cys Thr  
-5 1 5  
Glu Phe Met Ala Gly Leu Val Xaa Trp Leu Glu Leu Ser Glu Ala Val  
10 15 20 25  
Leu Pro Thr Met Thr Ala Phe Ala Ser Gly Leu Gly Gly Glu Gly Xaa  
30 35 40  
Xaa Cys Val Cys Ser Asn Phe Thr Glu Gly Pro His Leu Glu Gly Arg  
45 50 55  
Pro Asp Gly Asp His Ser Gly Pro Ser Glu Leu Leu Thr Gln Gly Trp  
60 65 70  
Ala Leu  
75

<210> 486  
<211> 209  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -84..-1

<400> 486  
Met Val Asn Phe Pro Gln Lys Ile Ala Gly Glu Leu Tyr Gly Pro Leu  
-80 -75 -70  
Met Leu Val Phe Thr Leu Val Ala Ile Leu Leu His Gly Met Lys Thr  
-65 -60 -55  
Ser Asp Thr Ile Ile Arg Glu Gly Thr Leu Met Gly Thr Ala Ile Gly  
-50 -45 -40  
Thr Cys Phe Gly Tyr Trp Leu Gly Val Ser Ser Phe Ile Tyr Phe Leu  
-35 -30 -25  
Ala Tyr Leu Cys Asn Ala Gln Ile Thr Met Leu Gln Met Leu Ala Leu  
-20 -15 -10 -5  
Leu Gly Tyr Gly Leu Phe Gly His Cys Ile Val Leu Phe Ile Thr Tyr  
1 5 10  
Asn Ile His Leu Arg Ala Leu Phe Tyr Leu Phe Trp Leu Leu Val Gly  
15 20 25  
Gly Leu Ser Thr Leu Arg Met Val Ala Val Leu Val Ser Arg Thr Val  
30 35 40  
Gly Pro Thr Xaa Arg Xaa Leu Leu Cys Gly Thr Leu Ala Ala Leu His

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```

45          50          55          60
Met Leu Phe Leu Leu Tyr Leu His Phe Ala Tyr His Lys Xaa Val Xaa
          65          70          75
Gly Ile Leu Asp Thr Leu Glu Gly Pro Asn Ile Pro Pro Ile Gln Arg
          80          85          90
Val Pro Arg Asp Ile Pro Ala Met Leu Pro Ala Ala Arg Leu Pro Thr
          95          100          105
Thr Val Leu Asn Ala Thr Ala Lys Ala Val Ala Val Thr Leu Gln Ser
          110          115          120
His
125
```

<210> 487  
<211> 36  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -17..-1

```

<400> 487
Met Gly Trp Gln Arg Trp Trp Cys Phe His Leu Gln Ala Glu Ala Ser
          -15          -10          -5
Ala His Pro Pro Gln Gly Leu Gln Ala Gln Phe Ser Cys Cys Pro Trp
          1          5          10          15
Val Gly Ile Cys
```

<210> 488  
<211> 44  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -29..-1

```

<400> 488
Met Met Ser Ser Glu Leu Arg Arg Asn Pro His Phe Leu Lys Ser Asn
          -25          -20          -15
Leu Phe Leu Gln Leu Leu Val Ser His Glu Ile Val Cys Ala Thr Glu
          -10          -5          1
Thr Val Thr Thr Asn Phe Leu Arg His Glu Lys Ala
          5          10          15
```

<210> 489  
<211> 163  
<212> PRT  
<213> Homo sapiens

<220>

<221> SIGNAL

<222> -52..-1

<400> 489

Met	Glu	His	Tyr	Arg	Lys	Ala	Gly	Ser	Val	Glu	Leu	Pro	Ala	Pro	Ser
	-50					-45						-40			
Pro	Met	Pro	Gln	Leu	Pro	Pro	Asp	Thr	Leu	Glu	Met	Arg	Val	Arg	Asp
	-35					-30					-25				
Gly	Ser	Lys	Ile	Arg	Asn	Leu	Leu	Gly	Leu	Ala	Leu	Gly	Arg	Leu	Glu
	-20				-15					-10					-5
Gly	Gly	Ser	Ala	Arg	His	Val	Val	Phe	Ser	Gly	Ser	Gly	Arg	Ala	Ala
			1				5						10		
Gly	Lys	Ala	Val	Ser	Cys	Ala	Glu	Ile	Val	Lys	Arg	Arg	Val	Pro	Gly
	15					20					25				
Leu	His	Gln	Leu	Thr	Lys	Leu	Xaa	Phe	Leu	Gln	Thr	Glu	Asp	Ser	Trp
	30				35					40					
Val	Pro	Xaa	Ser	Pro	Asp	Thr	Gly	Leu	Xaa	Pro	Leu	Thr	Val	Arg	Arg
	45				50					55					60
His	Val	Pro	Ala	Xaa	Trp	Val	Leu	Leu	Xaa	Arg	Asp	Pro	Leu	Asp	Pro
			65					70					75		
Asn	Glu	Cys	Gly	Tyr	Gln	Pro	Pro	Gly	Ala	Pro	Pro	Gly	Leu	Gly	Ser
			80					85					90		
Met	Pro	Ser	Ser	Ser	Cys	Gly	Pro	Arg	Ser	Xaa	Lys	Arg	Ala	Xaa	Xaa
		95					100					105			
Thr	Arg	Ser													

<210> 490

<211> 64

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -47..-1

<400> 490

Met	His	Gly	Phe	Glu	Ile	Ile	Ser	Leu	Lys	Glu	Glu	Ser	Pro	Leu	Gly
	-45						-40					-35			
Lys	Val	Ser	Gln	Gly	Pro	Leu	Phe	Asn	Val	Thr	Ser	Gly	Ser	Ser	Ser
	-30					-25					-20				
Pro	Val	Thr	Trp	Leu	Gly	Leu	Leu	Ser	Phe	Gln	Asn	Leu	His	Cys	Phe
	-15				-10					-5				1	
Pro	Asp	Leu	Pro	Thr	Glu	Met	Pro	Leu	Xaa	Ala	Lys	Gly	Xaa	Asn	Thr
		5					10						15		

<210> 491

<211> 218

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -50..-1

<400> 491

Met	His	His	Gly	Leu	Thr	Pro	Leu	Leu	Leu	Gly	Val	His	Glu	Gln	Lys
-50					-45					-40					-35
Gln	Gln	Val	Val	Lys	Phe	Leu	Ile	Lys	Lys	Lys	Ala	Asn	Leu	Asn	Ala
				-30					-25					-20	
Leu	Asp	Arg	Tyr	Gly	Arg	Thr	Ala	Leu	Ile	Leu	Ala	Val	Cys	Cys	Gly
			-15					-10					-5		
Ser	Ala	Ser	Ile	Val	Ser	Leu	Leu	Leu	Glu	Gln	Asn	Ile	Asp	Val	Ser
	1				5					10					
Ser	Gln	Asp	Leu	Ser	Gly	Gln	Thr	Ala	Lys	Lys	Tyr	Ala	Val	Ser	Ser
15				20					25					30	
Arg	His	Asn	Val	Ile	Cys	Gln	Leu	Leu	Ser	Asp	Tyr	Lys	Xaa	Lys	Gln
			35					40						45	
Xaa	Leu	Lys	Val	Ser	Ser	Glu	Asn	Ser	Asn	Pro	Xaa	Gln	Asp	Leu	Lys
		50					55					60			
Leu	Thr	Ser	Glu	Glu	Glu	Ser	Gln	Arg	Leu	Lys	Gly	Ser	Glu	Asn	Ser
	65					70						75			
Gln	Pro	Glu	Glu	Met	Ser	Gln	Glu	Pro	Glu	Ile	Asn	Xaa	Gly	Gly	Asp
80					85					90					
Arg	Lys	Val	Glu	Xaa	Xaa	Met	Lys	Lys	His	Gly	Ser	Xaa	His	Met	Gly
95				100						105					110
Phe	Pro	Xaa	Asn	Leu	Xaa	Asn	Gly	Ala	Thr	Ala	Asp	Asn	Gly	Asp	Asp
			115					120					125		
Gly	Leu	Ile	Pro	Pro	Xaa	Lys	Xaa	Xaa	Thr	Pro	Glu	Ser	Xaa	Gln	Phe
			130				135					140			
Pro	Asp	Thr	Glu	Asn	Glu	Gln	Tyr	His	Arg	Asp	Phe	Ser	Gly	His	Pro
	145				150						155				
Xaa	Phe	Pro	Thr	Thr	Leu	Pro	Ile	Lys	Gln						
	160				165										

<210> 492

<211> 216

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -15..-1

<400> 492

Met	Val	Cys	Val	Leu	Val	Leu	Ala	Ala	Ala	Ala	Gly	Ala	Val	Ala	Val
-15					-10					-5					1
Phe	Leu	Ile	Leu	Arg	Ile	Trp	Val	Val	Leu	Arg	Ser	Met	Asp	Val	Thr
		5					10					15			
Pro	Arg	Glu	Ser	Leu	Ser	Ile	Leu	Val	Val	Ala	Gly	Ser	Gly	Gly	His
	20				25							30			
Thr	Thr	Glu	Ile	Leu	Arg	Leu	Leu	Gly	Ser	Leu	Ser	Asn	Ala	Tyr	Ser
	35				40					45					
Pro	Arg	His	Tyr	Val	Ile	Ala	Asp	Thr	Asp	Glu	Met	Ser	Ala	Asn	Lys
50				55					60					65	
Ile	Asn	Ser	Phe	Glu	Leu	Xaa	Arg	Xaa	Asp	Arg	Xaa	Pro	Ser	Asn	Met
			70				75						80		
Xaa	Thr	Lys	Tyr	Tyr	Ile	His	Arg	Ile	Pro	Xaa	Ser	Arg	Glu	Val	Gln



85							90					95				
Gln	Ser	Trp	Pro	Ser	Thr	Val	Xaa	Thr	Thr	Leu	His	Ser	Met	Trp	Leu	
100							105					110				
Ser	Xaa	Pro	Leu	Ile	His	Arg	Val	Lys	Pro	Xaa	Leu	Val	Leu	Cys	Asn	
115							120					125				
Gly	Pro	Gly	Thr	Cys	Val	Pro	Ile	Cys	Val	Ser	Ala	Leu	Leu	Leu	Gly	
130	135					140					145					
Ile	Leu	Gly	Ile	Lys	Lys	Val	Ile	Ile	Val	Tyr	Val	Glu	Ser	Ile	Cys	
150							155					160				
Arg	Val	Lys	Thr	Leu	Ser	Met	Ser	Gly	Lys	Ile	Leu	Phe	His	Leu	Ser	
165							170					175				
Asn	Tyr	Phe	Ile	Val	Gln	Trp	Pro	Ala	Leu	Lys	Glu	Lys	Tyr	Pro	Lys	
180							185					190				
Ser	Val	Tyr	Leu	Gly	Arg	Ile	Val									
195							200									

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<210> 493
<211> 134
<212> PRT
<213> Homo sapiens
```

```

<220>
<221> SIGNAL
<222> -19..-1

```

```

4400> 493
Met Pro Leu Gly Ala Arg Ile Leu Phe His Gly Val Phe Tyr Ala Gly
      -15                      -10                      -5
Gly Phe Ala Ile Val Tyr Tyr Leu Ile Gln Lys Phe His Ser Arg Thr
      1                      5                      10
Leu Tyr Tyr Lys Leu Ala Val Glu Gln Leu Gln Xaa His Pro Glu Ala
      15                      20                      25
Gln Glu Ala Leu Gly Pro Pro Leu Asn Ile His Tyr Leu Lys Leu Ile
      30                      35                      40                      45
Asp Arg Glu Asn Phe Val Asp Ile Val Xaa Ala Lys Leu Lys Ile Pro
      50                      55                      60
Val Ser Gly Ser Lys Ser Glu Gly Leu Leu Tyr Val His Ser Ser Arg
      65                      70                      75
Gly Gly Pro Phe Gln Arg Trp His Leu Asp Glu Val Phe Leu Glu Leu
      80                      85                      90
Lys Asp Gly Gln Gln Ile Pro Val Phe Lys Leu Ser Gly Glu Asn Gly
      95                      100                      105
Asp Glu Val Lys Lys Glu
110                      115

```

```
<210> 494
<211> 85
<212> PRT
<213> Homo sapiens
```

```
<220>
<221> SIGNAL
<222> -16..-1
```

<400> 494

```

Met Ala Val Thr Ala Leu Ala Ala Xaa Thr Trp Leu Gly Val Trp Gly
  -15                -10                -5
Val Arg Thr Met Gln Ala Arg Gly Phe Gly Ser Asp Gln Ser Glu Asn
  1                5                10                15
Val Asp Arg Gly Ala Gly Ser Ile Arg Glu Ala Gly Gly Ala Phe Gly
                20                25                30
Lys Arg Glu Gln Ala Glu Glu Glu Arg Tyr Phe Arg Ala Gln Ser Thr
                35                40                45
Glu Gln Leu Ala Xaa Leu Lys Lys Xaa His Glu Glu Glu Ile Val His
                50                55                60
His Arg Glu Gly Asp
65

```

<210> 495

<211> 292

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -29...-1

<400> 495

```

Met His Gly Leu Leu His Tyr Leu Phe His Thr Arg Asn His Thr Phe
                -25                -20                -15
Ile Val Leu His Leu Val Leu Gln Gly Met Val Tyr Thr Glu Tyr Thr
                -10                -5                1
Trp Glu Val Phe Gly Tyr Cys Gln Glu Leu Glu Leu Ser Leu His Tyr
  5                10                15
Leu Leu Leu Pro Tyr Leu Leu Leu Gly Val Asn Leu Phe Phe Phe Thr
  20                25                30                35
Leu Thr Cys Gly Thr Asn Pro Gly Ile Ile Thr Lys Ala Asn Glu Leu
                40                45                50
Leu Phe Leu His Val Tyr Glu Phe Asp Glu Xaa Met Phe Pro Lys Asn
                55                60                65
Val Arg Cys Ser Thr Cys Asp Leu Arg Lys Pro Ala Arg Ser Xaa His
                70                75                80
Cys Xaa Val Cys Asn Trp Cys Val His Arg Phe Xaa His His Cys Val
  85                90                95
Trp Val Asn Asn Cys Ile Gly Ala Trp Asn Ile Arg Xaa Phe Leu Ile
  100                105                110                115
Tyr Val Leu Thr Leu Thr Ala Ser Ala Ala Thr Val Ala Ile Val Ser
                120                125                130
Thr Thr Phe Leu Val His Leu Val Val Met Ser Asp Leu Tyr Gln Glu
                135                140                145
Thr Tyr Ile Asp Asp Leu Gly His Leu His Val Met Asp Thr Val Phe
                150                155                160
Leu Ile Gln Tyr Leu Phe Leu Thr Phe Pro Arg Ile Val Phe Met Leu
                165                170                175
Gly Phe Val Val Val Leu Xaa Phe Leu Leu Gly Gly Tyr Leu Leu Phe
  180                185                190                195
Val Leu Tyr Leu Ala Ala Thr Asn Gln Thr Thr Asn Glu Trp Tyr Arg
                200                205                210

```

Xaa Asp Trp Ala Trp Cys Gln Arg Cys Pro Leu Val Ala Trp Pro Pro  
 215 220 225  
 Ser Ala Glu Pro Gln Val His Arg Asn Ile His Ser His Gly Leu Arg  
 230 235 240  
 Xaa Asn Leu Gln Glu Ile Phe Leu Pro Ala Phe Pro Cys His Glu Arg  
 245 250 255  
 Lys Lys Gln Glu  
 260

<210> 496  
 <211> 122  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -56..-1

<400> 496

Met Thr Gly Phe Leu Leu Pro Pro Ala Ser Arg Gly Thr Arg Arg Ser  
 -55 -50 -45  
 Cys Ser Arg Ser Arg Lys Arg Gln Thr Arg Arg Arg Arg Asn Pro Ser  
 40 -35 -30 -25  
 Ser Phe Val Ala Ser Cys Pro Thr Leu Leu Pro Phe Ala Cys Val Pro  
 -20 -15 -10  
 Gly Ala Ser Pro Thr Thr Leu Ala Phe Pro Pro Val Xaa Leu Thr Gly  
 -5 1 5  
 Pro Xaa Thr Asp Gly Ile Pro Phe Ala Leu Xaa Ser Ala Ala Gly Pro  
 10 15 20  
 Phe Cys Ala Ser Phe Pro Ser Gly Xaa Leu Ser Pro Pro Gly Pro Leu  
 25 30 35 40  
 Pro Gly Val Arg Gly Leu Pro Leu Pro Ser Val Phe Tyr Ser Cys Gly  
 45 50 55  
 Ala His Pro Lys Val Leu Lys Val Ala Leu  
 60 65

<210> 497  
 <211> 59  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -28..-1

<400> 497

Met Leu Xaa Leu Ser Arg Ala Thr Lys Xaa Gly Arg Ala Arg Trp Leu  
 -25 -20 -15  
 Met Pro Val Ile Pro Ala Leu Gln Glu Ala Xaa Ala Gly Gly Ser Arg  
 -10 -5 1  
 Gly Gln Glu Phe Glu Thr Ser Leu Ala Asn Met Glu Thr Glu Ala Gly  
 5 10 15 20  
 Glu Leu Leu Lys Pro Arg Arg Arg Arg Leu Gln

25

30

<210> 498  
<211> 99  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -13...-1

<400> 498  
Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg Arg Pro  
                  -10                  -5                  1  
Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu Ala His  
      5                  10                  15  
Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys Trp Arg  
20                  25                  30                  35  
Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn Ser Ser  
                  40                  45                  50  
Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr Pro Met  
                  55                  60                  65  
Arg Arg Ser Ser Cys His Leu Glu Cys Xaa Val Ile Phe Leu Leu Gly  
      70                  75                  80  
Arg Gln Leu  
      85

<210> 499  
<211> 99  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -13...-1

<400> 499  
Met His Leu Leu Ser Asn Trp Ala Asn Pro Ala Ser Ser Arg Arg Pro  
                  -10                  -5                  1  
Ser Met Ala Ala Ser Gly Thr Ser Trp Ile Ser Ser Thr Leu Ala His  
      5                  10                  15  
Ser Leu Ser Leu Arg Asp Val Ser Glu Arg Leu Cys Ser Cys Trp Arg  
20                  25                  30                  35  
Thr Ile Ser Met Gly Pro Cys Ala Arg Gly Ser Pro Met Asn Ser Ser  
                  40                  45                  50  
Gly Val His Arg Lys Ser Ser Arg Leu Phe Tyr Ile Arg Thr Pro Met  
                  55                  60                  65  
Arg Arg Ser Ser Cys His Leu Xaa Cys Gln Val Ile Phe Leu Leu Gly  
      70                  75                  80  
Arg Gln Leu  
      85

<210> 500  
<211> 108  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -25...-1

<400> 500  
Met Ser Leu Thr Ser Ser Ser Ser Val Arg Val Glu Trp Ile Ala Ala  
-25 -20 -15 -10  
Val Thr Ile Ala Ala Gly Thr Ala Ala Ile Gly Tyr Leu Ala Tyr Lys  
-5 1 5  
Arg Phe Tyr Val Lys Asp His Arg Asn Lys Ala Met Ile Asn Leu His  
10 15 20  
Ile Gln Lys Asp Asn Pro Lys Ile Val His Ala Phe Asp Met Glu Asp  
25 30 35  
Leu Gly Asp Lys Ala Val Tyr Cys Arg Cys Trp Arg Ser Lys Lys Phe  
40 45 50 55  
Pro Phe Cys Asp Gly Ala His Thr Lys His Asn Glu Glu Thr Gly Asp  
60 65 70  
Asn Val Gly Pro Leu Ile Ile Lys Lys Lys Glu Thr  
75 80

<210> 501  
<211> 183  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -15...-1

<400> 501  
Met Glu Ala Met Trp Leu Leu Cys Val Ala Leu Ala Val Leu Ala Trp  
-15 -10 -5 1  
Gly Phe Leu Trp Val Trp Asp Ser Ser Glu Arg Met Lys Ser Arg Glu  
5 10 15  
Gln Gly Arg Arg Leu Gly Ala Glu Ser Arg Thr Leu Leu Val Ile Ala  
20 25 30  
His Pro Asp Asp Glu Ala Met Phe Phe Ala Pro Thr Val Leu Gly Leu  
35 40 45  
Ala Arg Leu Arg His Trp Val Tyr Leu Leu Cys Phe Ser Ala Gly Asn  
50 55 60 65  
Tyr Tyr Asn Gln Gly Glu Thr Arg Lys Lys Glu Leu Leu Gln Ser Cys  
70 75 80  
Asp Val Leu Gly Ile Pro Leu Ser Ser Val Met Ile Ile Asp Asn Arg  
85 90 95  
Asp Phe Pro Xaa Asp Pro Gly Met Gln Trp Asp Thr Xaa His Val Ala  
100 105 110  
Xaa Val Leu Leu Gln His Ile Glu Val Asn Gly Ile Asn Leu Val Val  
115 120 125  
Thr Phe Asp Ala Gly Gly Xaa Ser Gly His Ser Asn His Ile Ala Leu

130		135		140		145									
Tyr	Ala	Ala	Val	Arg	Lys	Leu	Glu	Gly	Gln	Ile	Cys	Lys	Pro	Cys	Gly
		150						155						160	
Thr	Gly	Gln	Asp	Phe	Lys	Glu									
		165													

<210> 502  
 <211> 98  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -15..-1

<400> 502															
Met	Glu	Ala	Met	Trp	Leu	Leu	Cys	Val	Ala	Leu	Ala	Val	Leu	Ala	Trp
-15					-10					-5					1
Gly	Phe	Leu	Trp	Val	Trp	Asp	Ser	Ser	Glu	Arg	Met	Lys	Ser	Arg	Glu
		5					10					15			
Gln	Gly	Xaa	Arg	Leu	Gly	Ala	Glu	Ser	Arg	Thr	Leu	Leu	Val	Ile	Ala
		20				25					30				
His	Pro	Asp	Asp	Glu	Ala	Met	Phe	Phe	Ala	Pro	Thr	Val	Leu	Gly	Leu
		35				40					45				
Ala	Arg	Leu	Arg	His	Trp	Val	Tyr	Leu	Leu	Cys	Phe	Ser	Ala	Val	Phe
50					55					60					65
Arg	Arg	Glu	Leu	Ser	Glu	Tyr	Thr	Glu	Xaa	Leu	Thr	Ser	Glu	Pro	Leu
			70					75						80	
Xaa	Ala														

<210> 503  
 <211> 183  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -57..-1

<400> 503															
Met	Asp	Val	Thr	Gly	Asp	Glu	Glu	Glu	Glu	Ile	Lys	Gln	Glu	Ile	Asn
	-55					-50						-45			
Met	Leu	Lys	Lys	Tyr	Ser	His	His	Arg	Asn	Ile	Ala	Thr	Tyr	Tyr	Gly
	-40					-35					-30				
Ala	Phe	Ile	Lys	Lys	Asn	Pro	Pro	Gly	Met	Asp	Asp	Gln	Leu	Trp	Leu
-25					-20					-15					-10
Val	Met	Glu	Phe	Cys	Gly	Ala	Gly	Ser	Val	Thr	Asp	Leu	Ile	Lys	Asn
			-5					1				5			
Thr	Lys	Gly	Asn	Thr	Leu	Lys	Glu	Glu	Trp	Ile	Ala	Tyr	Ile	Cys	Xaa
	10					15					20				
Glu	Ile	Leu	Arg	Gly	Leu	Xaa	His	Leu	His	Gln	His	Lys	Val	Ile	His
	25				30				35						
Arg	Xaa	Ile	Lys	Gly	Gln	Asn	Val	Leu	Leu	Thr	Glu	Asn	Ala	Glu	Val

40					45					50					55
Lys	Leu	Val	Asp	Phe	Gly	Xaa	Xaa	Ala	Gln	Leu	Asp	Arg	Thr	Val	Gly
				60					65					70	
Arg	Xaa	Asn	Thr	Phe	Ile	Gly	Thr	Pro	Tyr	Trp	Met	Ala	Pro	Xaa	Val
			75					80					85		
Ile	Ala	Cys	Asp	Glu	Asn	Pro	Xaa	Ala	Thr	Tyr	Asp	Phe	Lys	Xaa	Asp
		90					95				100				
Leu	Trp	Ser	Leu	Gly	Ile	Thr	Ala	Ile	Glu	Met	Ala	Glu	Gly	Leu	Pro
	105					110					115				
Leu	Ser	Val	Thr	Cys	Thr	Pro									
120					125										

<210> 504  
<211> 140  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -14...-1

<400> 504															
Met	Phe	Leu	Thr	Ala	Leu	Leu	Trp	Arg	Gly	Arg	Ile	Pro	Gly	Arg	Gln
				-10					-5					1	
Trp	Ile	Gly	Lys	His	Arg	Arg	Pro	Arg	Phe	Val	Ser	Leu	Arg	Ala	Lys
		5					10					15			
Gln	Asn	Met	Ile	Arg	Arg	Leu	Glu	Ile	Glu	Ala	Glu	Asn	His	Tyr	Trp
	20					25					30				
Leu	Ser	Met	Pro	Tyr	Met	Thr	Arg	Glu	Gln	Glu	Arg	Gly	His	Ala	Ala
	35				40				45					50	
Leu	Arg	Arg	Arg	Glu	Ala	Phe	Glu	Ala	Ile	Lys	Ala	Ala	Ala	Thr	Ser
			55					60						65	
Lys	Phe	Pro	Pro	His	Arg	Phe	Ile	Ala	Asp	Gln	Leu	Asp	His	Leu	Asn
		70						75					80		
Xaa	His	Gln	Glu	Met	Val	Leu	Ile	Leu	Ser	Arg	His	Pro	Trp	Ile	Leu
	85						90					95			
Trp	Ile	Thr	Glu	Leu	Thr	Ile	Phe	Thr	Trp	Ser	Gly	Leu	Lys	Asn	Cys
	100					105					110				
Ser	Leu	Cys	Glu	Asn	Glu	Leu	Trp	Thr	Ser	Leu	Tyr				
115					120					125					

<210> 505  
<211> 59  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -14...-1

<400> 505  
Met Ala Ala Leu Val Thr Val Leu Phe Thr Gly Val Arg Arg Leu His  
-10 -5 1

Cys	Ser	Ala	Xaa	Leu	Gly	Arg	Ala	Ala	Ser	Gly	Xaa	Tyr	Ser	Arg	Asn
	5						10					15			
Trp	Leu	Pro	Thr	Pro	Pro	Ala	Thr	Gly	Pro	Leu	Pro	Ser	Ser	Gln	Thr
	20					25					30				
Gly	His	Met	Arg	Met	Ala	Ala	Leu	Leu	Pro	Gln					
35					40					45					

<210> 506  
 <211> 101  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -36...-1

Met	Gly	Pro	Tyr	Asn	Val	Ala	Val	Pro	Ser	Asp	Val	Ser	His	Ala	Arg
	-35					-30					-25				
Phe	Tyr	Phe	Leu	Phe	His	Arg	Pro	Leu	Arg	Leu	Leu	Asn	Leu	Leu	Ile
	20				-15					-10					-5
Leu	Ile	Glu	Gly	Ser	Val	Val	Phe	Tyr	Gln	Leu	Tyr	Ser	Leu	Leu	Arg
				1				5					10		
Ser	Glu	Lys	Trp	Asn	His	Thr	Leu	Ser	Met	Ala	Leu	Ile	Leu	Phe	Cys
		15					20					25			
Asn	Tyr	Tyr	Val	Leu	Phe	Lys	Leu	Leu	Arg	Asp	Arg	Xaa	Xaa	Leu	Gly
	30					35				40					
Arg	Ala	Tyr	Ser	Tyr	Pro	Leu	Asn	Ser	Tyr	Glu	Leu	Lys	Ala	Asn	Xaa
	45				50					55					60
Ala	Ala	Ser	Xaa	Gln											
				65											

<210> 507  
 <211> 341  
 <212> PRT  
 <213> Homo sapiens

<220>  
 <221> SIGNAL  
 <222> -55...-1

Met	Arg	Lys	Val	Val	Leu	Ile	Thr	Gly	Ala	Ser	Ser	Gly	Ile	Gly	Leu
	-55				-50					-45					-40
Ala	Leu	Cys	Lys	Arg	Leu	Leu	Ala	Glu	Asp	Asp	Glu	Leu	His	Leu	Cys
			-35						-30					-25	
Leu	Ala	Cys	Arg	Asn	Met	Ser	Lys	Ala	Glu	Ala	Val	Cys	Ala	Ala	Leu
			-20					-15					-10		
Leu	Ala	Ser	His	Pro	Thr	Ala	Glu	Val	Thr	Ile	Val	Gln	Val	Asp	Val
	-5						1				5				
Ser	Asn	Leu	Gln	Ser	Phe	Phe	Arg	Ala	Ser	Lys	Glu	Leu	Lys	Gln	Arg
10					15					20					25
Phe	Gln	Arg	Leu	Asp	Cys	Ile	Tyr	Leu	Asn	Ala	Gly	Ile	Met	Pro	Asn



Pro	Gln	Leu	Asn	Ile	Lys	Ala	Leu	Phe	Phe	Gly	Leu	Phe	Ser	Arg	Lys
			45					50					55		
Val	Ile	His	Met	Phe	Ser	Thr	Ala	Glu	Gly	Leu	Leu	Thr	Gln	Gly	Asp
		60					65					70			
Lys	Ile	Thr	Ala	Asp	Gly	Leu	Gln	Glu	Val	Phe	Glu	Thr	Asn	Val	Phe
	75					80				85					
Gly	His	Phe	Ile	Leu	Ile	Arg	Glu	Leu	Glu	Pro	Leu	Leu	Cys	His	Ser
90					95					100					105
Asp	Asn	Pro	Ser	Gln	Leu	Ile	Trp	Thr	Ser	Ser	Arg	Ser	Ala	Arg	Lys
				110					115					120	
Ser	Asn	Phe	Ser	Leu	Glu	Asp	Phe	Gln	His	Ser	Lys	Gly	Lys	Glu	Pro
			125					130					135		
Tyr	Ser	Ser	Ser	Lys	Tyr	Ala	Thr	Asp	Leu	Leu	Ser	Val	Ala	Leu	Asn
		140					145					150			
Arg	Asn	Phe	Asn	Gln	Gln	Gly	Leu	Tyr	Ser	Asn	Val	Ala	Cys	Pro	Gly
	155					160					165				
Thr	Ala	Leu	Thr	Asn	Leu	Thr	Tyr	Gly	Ile	Leu	Pro	Pro	Phe	Ile	Trp
170					175					180					185
Thr	Leu	Leu	Met	Pro	Ala	Ile	Leu	Leu	Leu	Arg	Phe	Phe	Ala	Asn	Ala
				190					195					200	
Phe	Thr	Leu	Thr	Pro	Tyr	Asn	Gly	Thr	Glu	Ala	Leu	Val	Trp	Leu	Phe
			205					210					215		
His	Gln	Lys	Pro	Glu	Ser	Leu	Asn	Pro	Leu	Ile	Lys	Tyr	Leu	Ser	Ala
		220					225					230			
Thr	Thr	Gly	Phe	Gly	Arg	Asn	Tyr	Ile	Met	Thr	Gln	Lys	Met	Asp	Leu
	235					240					245				
Asp	Glu	Asp	Thr	Ala	Glu	Lys	Phe	Tyr	Gln	Lys	Leu	Leu	Glu	Leu	Glu
250					255					260					265
Lys	His	Ile	Arg	Val	Thr	Ile	Gln	Lys	Thr	Asp	Asn	Gln	Ala	Arg	Leu
				270					275					280	
Ser	Gly	Ser	Cys	Leu											
			285												

<210> 508

<211> 108

<212> PRT

<213> Homo sapiens

<220>

<221> SIGNAL

<222> -42..-1

<400> 508

Met	His	Ile	Leu	Gln	Leu	Leu	Thr	Thr	Val	Asp	Asp	Gly	Ile	Gln	Ala
			-40				-35					-30			
Ile	Val	His	Cys	Pro	Asp	Thr	Gly	Lys	Asp	Ile	Trp	Asn	Leu	Leu	Phe
		-25				-20				-15					
Asp	Leu	Val	Cys	His	Glu	Phe	Cys	Gln	Ser	Asp	Asp	Pro	Ala	Ile	Ile
-10					-5				1					5	
Leu	Gln	Xaa	Gln	Lys	Thr	Val	Leu	Ala	Ser	Val	Phe	Ser	Val	Leu	Ser
			10					15					20		
Ala	Ile	Tyr	Ala	Ser	Gln	Thr	Glu	Gln	Xaa	Tyr	Leu	Lys	Ile	Xaa	Lys
		25					30					35			
Gly	Asp	Gly	Gly	Ser	Gly	Ser	Lys	Gly	Arg	Pro	Xaa	Xaa	Gln	Thr	Glu

40						45						50
Xaa	Phe	Leu	Cys	Ile	Ser	Lys	Pro	Ser	Ser	Phe	Leu	
55					60					65		

<210> 509  
<211> 80  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -26...-1

<400> 509  
Met Glu Glu Ile Ser Ser Pro Leu Val Glu Phe Val Lys Val Leu Cys  
-25 -20 -15  
Thr Asn Gln Val Leu Ile Thr Ala Arg Ala Val Pro Thr Lys Lys Ala  
-10 -5 1 5  
Ser Val Arg Cys Val Glu Lys Arg Phe Trp Ile Pro Lys Thr Thr Ser  
10 15 20  
Lys His Leu Ser Arg Cys Ile Asp Gly Ile Ser Gly Phe Leu Asn Asp  
25 30 35  
Phe Thr Phe Cys Leu Glu Phe Ser Arg His Arg Cys Gln Leu Thr Glu  
40 45 50

<210> 510  
<211> 158  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -44...-1

<400> 510  
Met Ala Gly Phe Leu Asp Asn Phe Arg Trp Pro Glu Cys Glu Cys Ile  
-40 -35 -30  
Asp Trp Ser Glu Arg Arg Asn Ala Val Ala Ser Val Val Ala Gly Ile  
-25 -20 -15  
Leu Phe Phe Thr Gly Trp Trp Ile Met Ile Asp Ala Ala Val Val Tyr  
-10 -5 1  
Pro Lys Pro Glu Gln Leu Asn His Ala Phe His Thr Cys Gly Val Phe  
5 10 15 20  
Ser Thr Leu Ala Phe Phe Met Ile Asn Ala Val Ser Asn Ala Gln Val  
25 30 35  
Arg Gly Asp Ser Tyr Glu Ser Gly Cys Leu Gly Arg Thr Gly Ala Arg  
40 45 50  
Val Trp Leu Phe Ile Gly Phe Met Leu Met Phe Gly Ser Leu Ile Ala  
55 60 65  
Ser Met Trp Ile Leu Phe Gly Ala Tyr Val Thr Gln Asn Thr Asp Val  
70 75 80  
Tyr Pro Gly Leu Ala Val Phe Phe Gln Asn Ala Leu Ile Phe Phe Ser  
85 90 95 100

Thr Leu Ile Tyr Lys Phe Gly Arg Thr Glu Glu Leu Trp Thr  
105 110

<210> 511  
<211> 130  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -28...-1

<400> 511  
Met Asn Trp Glu Leu Leu Leu Trp Leu Leu Val Leu Cys Ala Leu Leu  
-25 -20 -15  
Leu Leu Leu Val Gln Leu Leu Arg Phe Leu Arg Ala Asp Gly Asp Leu  
-10 -5 1  
Thr Leu Leu Trp Ala Glu Trp Gln Gly Arg Arg Pro Glu Trp Glu Leu  
5 10 15 20  
Thr Asp Met Val Val Trp Val Thr Gly Ala Ser Ser Gly Ile Gly Glu  
25 30 35  
Glu Leu Ala Tyr Gln Leu Ser Lys Leu Gly Val Ser Leu Val Leu Ser  
40 45 50  
Ala Arg Arg Val His Glu Leu Glu Arg Val Lys Arg Arg Cys Leu Glu  
55 60 65  
Asn Gly Asn Leu Lys Glu Lys Asp Ile Leu Val Leu Pro Leu Asp Leu  
70 75 80  
Thr Asp Thr Gly Ser His Glu Ser Gly Tyr Gln Ser Cys Ser Pro Gly  
85 90 95 100  
Ile Trp

<210> 512  
<211> 199  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -62...-1

<400> 512  
Met Ser Gln Arg Ser Leu Cys Met Asp Thr Ser Leu Asp Val Tyr Arg  
-60 -55 -50  
Xaa Leu Ile Glu Leu Asn Tyr Leu Gly Thr Val Ser Leu Thr Lys Cys  
-45 -40 -35  
Val Leu Pro His Met Ile Glu Arg Lys Gln Gly Lys Ile Val Thr Val  
-30 -25 -20 -15  
Asn Ser Ile Leu Gly Ile Ile Ser Val Pro Leu Ser Ile Gly Tyr Cys  
-10 -5 1  
Ala Ser Lys His Ala Leu Arg Gly Phe Phe Asn Gly Leu Arg Thr Glu  
5 10 15  
Leu Ala Thr Tyr Pro Gly Ile Ile Val Ser Asn Ile Cys Pro Gly Pro  
20 25 30

Val	Gln	Ser	Asn	Ile	Val	Glu	Asn	Ser	Leu	Ala	Gly	Glu	Val	Thr	Lys
35					40					45					50
Thr	Ile	Gly	Asn	Asn	Gly	Asn	Gln	Ser	His	Lys	Met	Thr	Thr	Ser	Arg
			55						60					65	
Cys	Val	Arg	Leu	Met	Leu	Ile	Ser	Met	Ala	Asn	Asp	Leu	Lys	Glu	Val
			70					75					80		
Trp	Ile	Ser	Glu	Gln	Pro	Phe	Leu	Leu	Val	Thr	Tyr	Leu	Trp	Gln	Tyr
		85					90					95			
Met	Pro	Thr	Trp	Ala	Trp	Trp	Ile	Thr	Asn	Lys	Met	Gly	Lys	Lys	Arg
	100					105					110				
Ile	Glu	Asn	Phe	Lys	Ser	Gly	Val	Asp	Ala	Xaa	Ser	Ser	Tyr	Phe	Lys
115					120					125					130
Ile	Phe	Lys	Thr	Lys	His	Asp									
				135											

<210> 513  
<211> 180  
<212> PRT  
<213> Homo sapiens

<220>  
<221> SIGNAL  
<222> -25..-1

Met	Asn	Thr	Val	Leu	Ser	Arg	Ala	Asn	Ser	Leu	Phe	Ala	Phe	Ser	Leu
-25					-20					-15					-10
Ser	Val	Met	Ala	Ala	Leu	Thr	Phe	Gly	Cys	Phe	Ile	Xaa	Thr	Ala	Phe
			-5					1				5			
Lys	Asp	Arg	Ser	Val	Pro	Val	Arg	Leu	His	Val	Ser	Arg	Ile	Met	Leu
	10					15					20				
Lys	Asn	Val	Glu	Asp	Phe	Thr	Gly	Pro	Arg	Glu	Arg	Ser	Asp	Leu	Gly
	25					30				35					
Phe	Ile	Thr	Phe	Asp	Ile	Thr	Ala	Asp	Leu	Glu	Asn	Ile	Phe	Asp	Trp
40					45					50					55
Asn	Val	Lys	Gln	Leu	Phe	Leu	Tyr	Leu	Ser	Ala	Glu	Tyr	Ser	Thr	Lys
			60					65					70		
Asn	Asn	Ala	Leu	Asn	Gln	Xaa	Val	Leu	Trp	Asp	Lys	Ile	Val	Leu	Arg
		75						80					85		
Gly	Asp	Asn	Pro	Lys	Leu	Leu	Leu	Lys	Asp	Met	Lys	Thr	Lys	Tyr	Phe
	90					95						100			
Phe	Phe	Asp	Asp	Gly	Asn	Gly	Leu	Xaa	Gly	Asn	Arg	Asn	Val	Thr	Leu
	105					110					115				
Thr	Leu	Ser	Trp	Asn	Val	Val	Pro	Asn	Ala	Gly	Ile	Leu	Pro	Leu	Val
120					125					130					135
Thr	Gly	Ser	Gly	His	Val	Ser	Val	Pro	Phe	Pro	Asp	Thr	Tyr	Glu	Ile
			140						145					150	
Thr	Lys	Ser	Tyr												
			155												

<210> 514  
<211> 120  
<212> PRT

<213> Bos taurus

<400> 514

Met	Met	Thr	Gly	Arg	Gln	Gly	Arg	Ala	Thr	Phe	Gln	Phe	Leu	Pro	Asp
1				5				10					15		
Glu	Ala	Arg	Ser	Leu	Pro	Pro	Pro	Lys	Leu	Thr	Asp	Pro	Arg	Leu	Ala
			20					25				30			
Phe	Val	Gly	Phe	Leu	Gly	Tyr	Cys	Ser	Gly	Leu	Ile	Asp	Asn	Ala	Ile
			35				40					45			
Arg	Arg	Arg	Pro	Val	Leu	Leu	Ala	Gly	Leu	His	Arg	Gln	Leu	Leu	Tyr
			50				55				60				
Ile	Thr	Ser	Phe	Val	Phe	Val	Gly	Tyr	Tyr	Leu	Leu	Lys	Arg	Gln	Asp
65				70						75				80	
Tyr	Met	Tyr	Ala	Val	Arg	Asp	His	Asp	Met	Phe	Ser	Tyr	Ile	Lys	Ser
				85				90					95		
His	Pro	Glu	Asp	Phe	Pro	Glu	Lys	Asp	Lys	Lys	Thr	Tyr	Gly	Glu	Val
			100					105					110		
Phe	Glu	Glu	Phe	His	Pro	Val	Arg								
			115				120								

<210> 515

<211> 1082

<212> DNA

<213> Homo sapiens

<400> 515

gatcccagac	ctcggcttgc	agtagtggtta	gactgaagat	aaagtaagtg	ctgtttgggc	60
taacaggatc	tcctcttgca	gtctgcagcc	caggacgctg	attccagcag	cgccttaccg	120
cgcagcccga	agattcacta	tggtgaaaat	cgccttcaat	acccctaccg	ccgtgcaaaa	180
ggaggaggcg	cggcaagacg	tggaggccct	cctgagccgc	acggtcagaa	ctcagatact	240
gaccggcaag	gagctccgag	ttgccaccca	ggaaaaagag	ggctcctctg	ggagatgtat	300
gcttactctc	ttaggccttt	cattcatctt	ggcaggactt	attgttggtg	gagcctgcat	360
ttacaagtac	ttcatgcccc	agagcaccat	ttaccgtgga	gagatgtgct	tttttgattc	420
tgaggatcct	gcaaattccc	ttcgtggagg	agagcctaac	ttcctgcctg	tgactgagga	480
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## EXTENDED cDNAs

### Abstract of the Disclosure

The sequences of extended cDNAs encoding secreted proteins are disclosed. The extended cDNAs can be used to express secreted proteins or portions thereof or to obtain antibodies capable of specifically binding to the secreted proteins. The extended cDNAs may also be used in diagnostic, forensic, gene therapy, and chromosome mapping procedures. The extended cDNAs may also be used to design expression vectors and secretion vectors.

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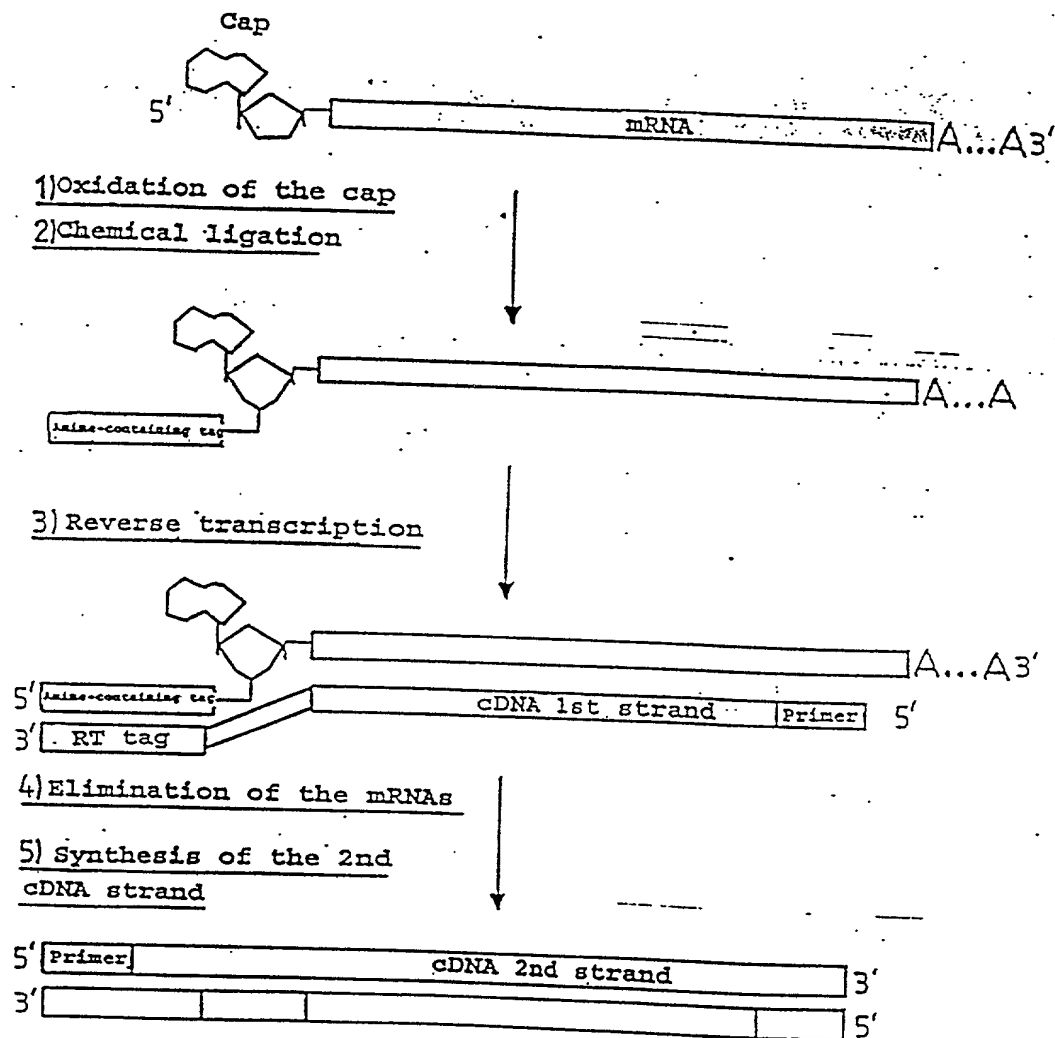


FIGURE 1

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Minimum signal peptide score	false positive rate	false negative rate	proba(0.1)	proba(0.2)
3,5	0,121	0,036	0,467	0,664
4	0,096	0,06	0,519	0,708
4,5	0,078	0,079	0,565	0,745
5	0,062	0,098	0,615	0,782
5,5	0,05	0,127	0,659	0,813
6	0,04	0,163	0,694	0,838
6,5	0,033	0,202	0,725	0,855
7	0,025	0,248	0,763	0,878
7,5	0,021	0,304	0,78	0,889
8	0,015	0,368	0,816	0,909
8,5	0,012	0,418	0,836	0,92
9	0,009	0,512	0,856	0,93
9,5	0,007	0,581	0,863	0,934
10	0,006	0,679	0,835	0,918

FIGURE 2

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Score curves

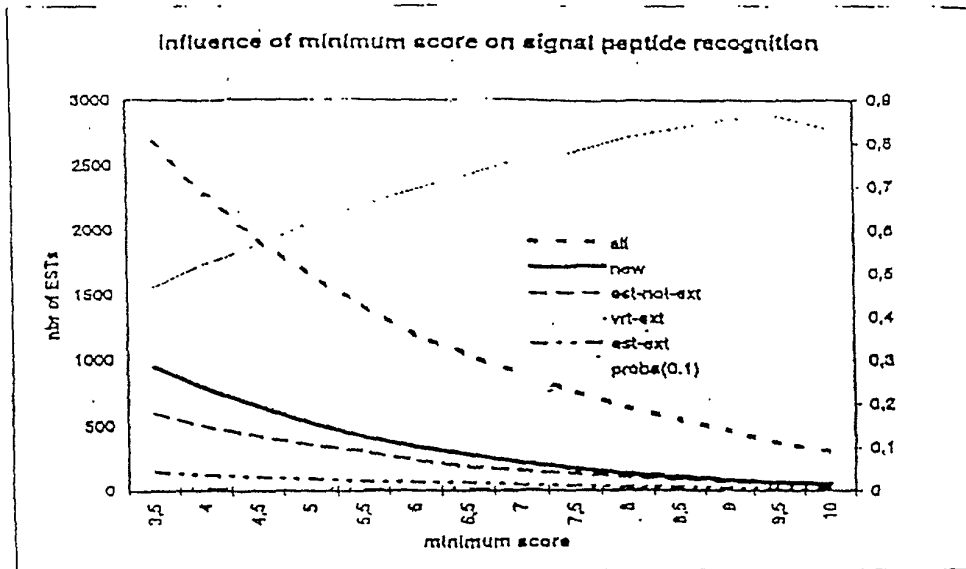


FIGURE 3

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Minimum signal peptide score	All ESTs	New ESTs	ESTs matching public EST closer than 40 bp from beginning	ESTs extending known mRNA more than 40 bp	ESTs extending public EST more than 40 bp
3,5	2874	947	599	23	150
4	2278	784	488	23	126
4,5	1943	647	425	22	112
5	1657	523	353	21	96
5,5	1417	419	307	19	80
6	1190	340	238	18	68
6,5	1035	280	186	18	60
7	893	219	161	15	48
7,5	753	173	132	12	36
8	636	133	101	11	29
8,5	643	104	83	8	26
9	456	81	63	6	24
9,5	364	57	48	6	18
10	303	47	35	6	15

FIGURE 4

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Tissue	All ESTs	New ESTs	ESTs matching public EST closer than 40 bp from beginning	ESTs extending known mRNA more than 40 bp	ESTs extending public EST more than 40 bp
Brain	329	131	75	3	24
Cancerous prostate	134	40	37	1	6
Cerebellum	17	9	1	0	6
Colon	21	11	4	0	0
Dystrophic muscle	41	18	8	0	1
Fetal brain	70	37	16	0	1
Fetal kidney	227	116	46	1	19
Fetal liver	13	7	2	0	0
Heart	30	15	7	0	1
Hypertrophic prostate	86	23	22	2	2
Kidney	10	7	3	0	0
Large intestine	21	8	4	0	1
Liver	23	9	6	0	0
Lung	24	12	4	0	1
Lung (cells)	57	38	6	0	4
Lymph ganglia	163	60	23	2	12
Lymphocytes	23	6	4	0	2
Muscle	33	16	6	0	4
Normal prostate	181	61	45	7	11
Ovary	90	57	12	1	2
Pancreas	48	11	6	0	1
Placenta	24	5	1	0	0
Prostate	34	16	4	0	2
Spleen	56	28	10	0	1
Substantia nigra	108	47	27	1	6
Testis	15	3	3	1	0
Thyroid	131	68	25	1	8
Umbilical cord	17	8	2	0	2
Uterus	55	17	12	1	3
Non tissue-specific	28	15	3	0	2
Total	568	48	177	2	28
	2677	947	601	23	150

FIGURE 5

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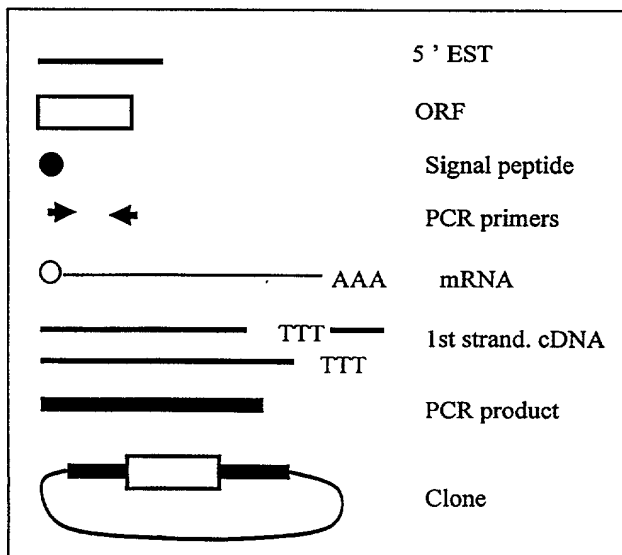
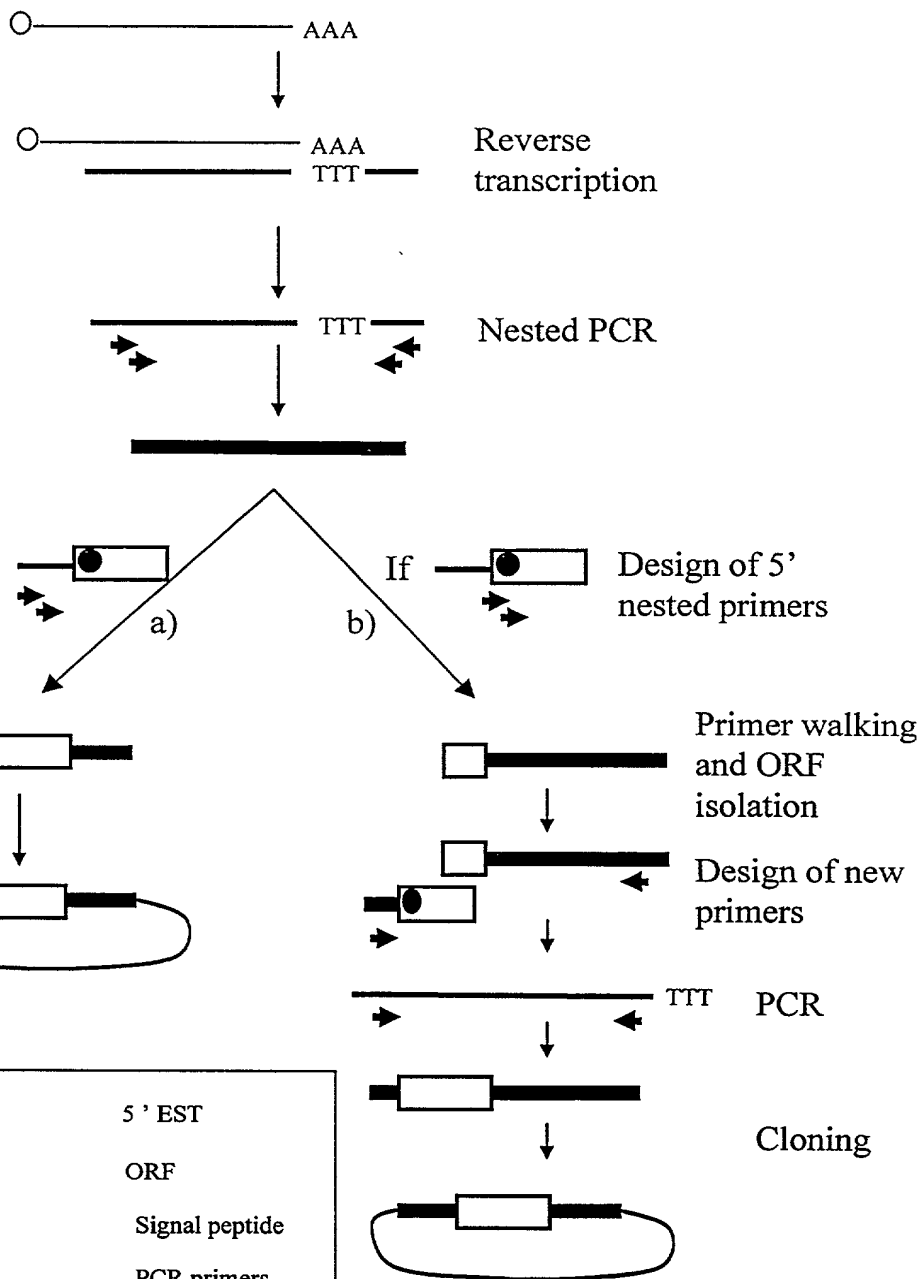
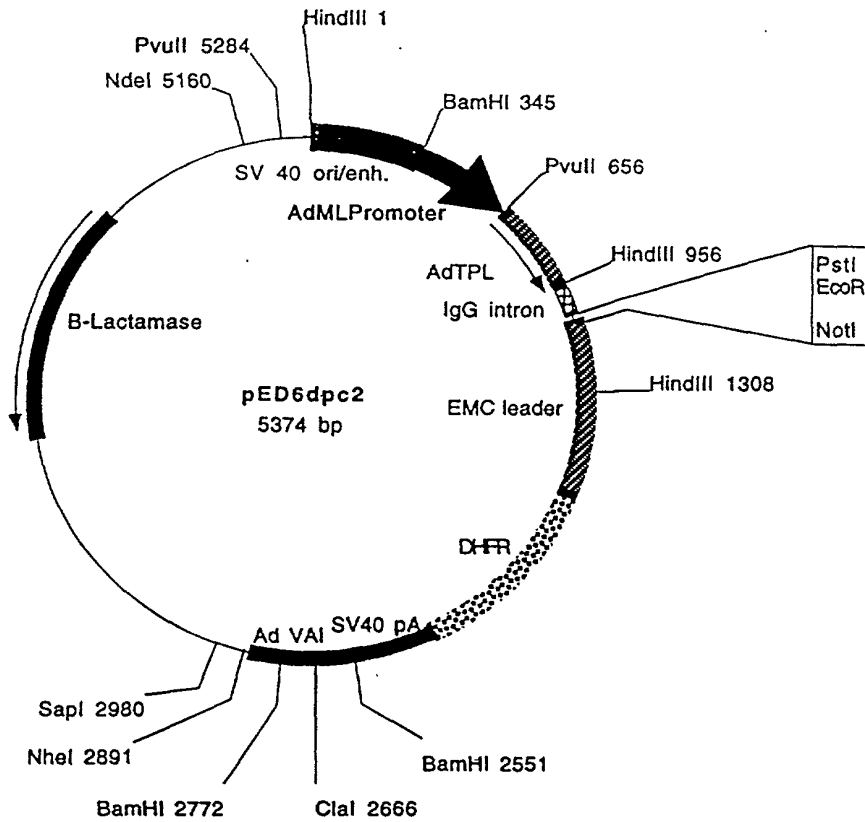


Figure 6

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Plasmid name: pED6dpc2

Plasmid size: 5374 bp

Comments/References: pED6dpc2 is derived from pED6dpc1 by insertion of a new polylinker to facilitate cDNA cloning. SST cDNAs are cloned between EcoRI and NotI. pED vectors are described in Kaufman et al.(1991), NAR 19: 4485-4490.

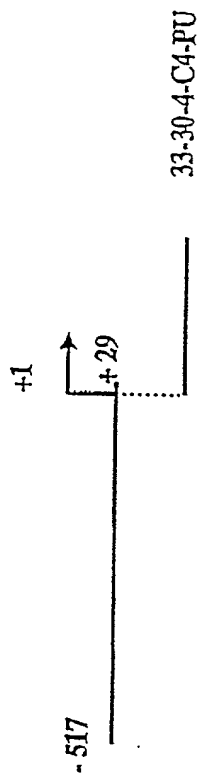
FIGURE 7



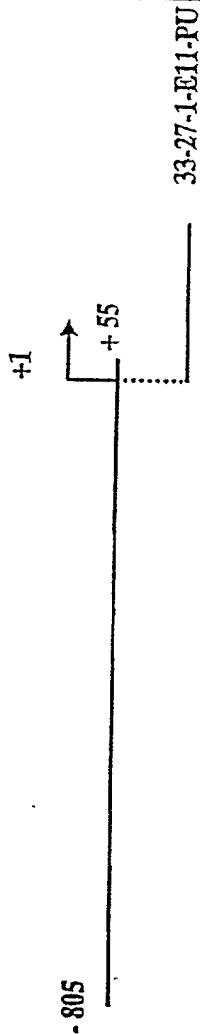
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Description of Promoter structure isolated from SignalTag 5' ESTs

Promoter P13H2



Promoter P15B4



Promoter P29B6

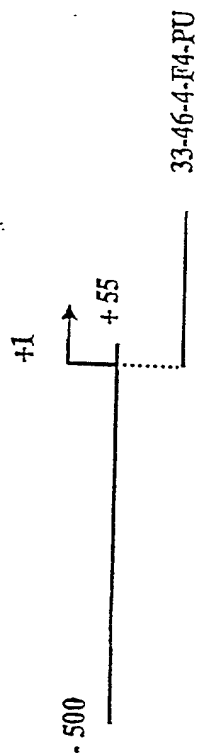


FIGURE 8

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Description of Transcription Factor Binding Sites present on promoters isolated from SignalTag sequences.

Promoter sequence P1342 (548 bp):

Matrix	Position	Orientation	Score	Length	Sequence
CMYB_01	-502	+	0.983	9	TGTCAGTTG
MYOQ_Q8	-501	-	0.981	10	CCCAACTGAC
S8_01	-444	-	0.980	11	AATAGAATTAG
S8_01	-425	+	0.988	11	AAGTAAATTAG
DELTAEF1_01	-390	-	0.980	11	GCACACOTCAG
GATA_C	-384	-	0.984	11	AGATAAATCCA
CMYB_01	-348	+	0.958	9	CTTCAGTTG
GATA1_02	-343	+	0.959	14	TTGTAGATAGGACA
GATA_C	-339	+	0.953	11	AGATAGGACAT
TAL1ALPHA47_01	-235	+	0.973	18	CATAACAGATGGTAAG
TAL1BETA47_01	-235	+	0.983	16	CATAACAGATGGTAAG
TAL1BETA172_01	-235	+	0.978	16	CATAACAGATGGTAAG
MYOQ_Q8	-232	-	0.954	10	ACCATCTGTT
GATA1_04	-217	-	0.953	13	TCAAGATAAAGTA
IK1_01	-128	+	0.963	13	AGTTGGGAATTCC
IK2_01	-128	+	0.985	12	AGTTGGGAATTCC
CREL_01	-123	+	0.982	10	TGGGAATTCC
GATA1_02	-98	+	0.980	14	TCAGTGATATGGCA
SRY_Q2	-41	-	0.951	12	TAAACAAAACA
E2F_Q2	-33	+	0.957	8	TTTAGCGC
MZF1_01	-6	-	0.976	8	TGAGGGGA

Promoter sequence P1684 (881 bp):

Matrix	Position	Orientation	Score	Length	Sequence
NFY_Q8	-748	-	0.958	11	GGACCAATCAT
MZF1_01	-738	+	0.962	8	CCTGGGGA
CMYB_01	-684	+	0.994	9	TGACCGTTG
VMYB_02	-682	-	0.985	9	TCCAACGGT
STAT_01	-673	+	0.988	9	TTCTGGAA
STAT_01	-673	-	0.951	9	TTCCAGGAA
MZF1_01	-656	-	0.958	8	TTGGGGGA
IK2_01	-451	+	0.985	12	GAATGGGATTTC
MZF1_01	-424	+	0.988	8	AGAGGGGA
SRY_Q2	-388	-	0.955	12	GAAAAAACAACA
MZF1_01	-216	+	0.960	8	GAAGGGGA
MYOQ_Q8	-190	+	0.981	10	AGCATCTGCC
DELTAEF1_01	-178	+	0.958	11	TCCCACCTTC
S8_01	6	-	0.982	11	GAGGCAATTAT
MZF1_01	16	-	0.988	8	AGAGGGGA

Promoter sequence P2986 (555 bp):

Matrix	Position	Orientation	Score	Length	Sequence
ARNT_01	-311	+	0.984	16	GGACTCACGTGCTGCT
NMYC_01	-309	+	0.985	12	ACTCACGTGCTG
USF_01	-309	+	0.985	12	ACTCACGTGCTG
USF_01	-309	-	0.985	12	CAGCAOGTGAGT
NMYC_01	-309	-	0.956	12	CAGCAOGTGAGT
MYCMA_02	-309	-	0.972	12	CAGCAOGTGAGT
USF_C	-307	+	0.997	8	TCACGTGC
USF_C	-307	-	0.991	8	GCACGTGA
MZF1_01	-292	-	0.988	8	CATGGGGA
ELK1_02	-105	+	0.963	14	CTCTCCGGAAGCCT
CETBP4_01	-102	+	0.974	10	TCCGGAAGCC
AP1_Q4	-42	-	0.963	11	AGTGACTGAAC
AP1FJ_Q2	-42	-	0.981	11	AGTGACTGAAC
PADS_C	45	+	1.000	9	TGTGGTCTC

Figure 9

100.0% identity in 125 aa overlap

```

      10      20      30      40      50      60
SEQ ID NO: 217 MADEELEALRRQRLAELQAKHGDPGDAAQQEAKHREAEMRNSILAQVLDQSARARLSNLA
      X::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::
SEQ ID NO: 516 MADEELEALRRQRLAELQAKHGDPGDAAQQEAKHREAEMRNSILAQVLDQSARARLSNLA
      10      20      30      40      50      60

      70      80      90     100     110     120
SEQ ID NO: 217 LVKPEKTKAVENYLIQMARYGQLSEKVSEQGLIEILKKVSQQTEKTTTVKFNRRKVMDS
      ::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::::
SEQ ID NO: 516 LVKPEKTKAVENYLIQMARYGQLSEKVSEQGLIEILKKVSQQTEKTTTVKFNRRKVMDS
      70      80      90     100     110     120

```

SEQ ID NO: 217 EDDDY

::::X

SEQ ID NO: 516 EDDDY

-----

**FIGURE 10**

## CLUSTAL W(1.5) multiple sequence alignment

```

SEQ ID NO: 517      MFCPLKLILLPVLLDYSLSGLNDLNVSPPELTVHVGDSALMGCVFQSTEDKCIFKIDWTLS
SEQ ID NO: 232      -----MGCVFQSTEDKCIFKIDWTLS
SEQ ID NO: 174      -----MGCVFQSTEDKRIFKIDWTLS
SEQ ID NO: 175      -----MGCVFQSTVDKCIFKIDWTLS
                      ***** ** *****

```

```

SEQ ID NO: 517      PGEHAKDEYVLYYYSNLSVPIGRFQNRVHLMGDNLCNDGSLLLQDVQDVE-----
SEQ ID NO: 232      PGEHAKDEYVLYYYSNLSVPIGRFQNRVHLMGDILCNDGSLLLQDVQEADQGTYYICEIRL
SEQ ID NO: 174      PGEHAKDEYVLYYYSNLSVPIGRFQNRVHLMGDNLCNDGSLLLQDVQEADQGTYYICEIRL
SEQ ID NO: 175      PGEHAKDEYVLYYYSNLSVPIGRFQNRVHLMGDILCNDGSLLLQDVQEADQGTYYICEIRL
                      *****

```

```

SEQ ID NO: 517      -----
SEQ ID NO: 232      KGESQVFKKAVVLHVLPEEPKGTQMLT-----
SEQ ID NO: 174      KGESQVFKKAVVLHVLPEEPKELMVHVGGLIQMGCVFQSTEVKHVTKVEWIFSGRRAKEE
SEQ ID NO: 175      KGESQVFKKAVVLHVLPEEPKELMVHVGGLIQMGCVFQSTEVKHVTKVEWIFSGR--RAK

```

```

SEQ ID NO: 517      -----
SEQ ID NO: 232      -----
SEQ ID NO: 174      IVFRYYHKLMSAEYSQSWGHFQNRVNLVGDI FRNDGSIMLQGVRESGGNYTCSIHLGN
SEQ ID NO: 175      VTRRKHHCVREGSG-----

```

```

SEQ ID NO: 517      -----
SEQ ID NO: 232      -----
SEQ ID NO: 174      LVFKKTIVLHVSPEEPRTLVTTPAALRPLVLGGNQLVIVGIVCATILLLPVLILIVKKTCTC
SEQ ID NO: 175      -----

```

```

SEQ ID NO: 517      -----
SEQ ID NO: 232      -----
SEQ ID NO: 174      GNKSSVNSTVLVKNTKKTNP
SEQ ID NO: 175      -----

```

FIGURE 11

		10	20	30	40	50	60	
SEQ ID NO: 515	PTAVQKEEARQDVEALLSRTVRTQILTGKELRVATQ	KEGSSGR	CMLTLLGLS	FILAGLI				
SEQ ID NO: 231					LRVATQ	KEGSSGR	CMLTLLGLS	FILAGLI
					10	20	30	
	70	80	90	100	110	120		
SEQ ID NO: 515	VGGACIYKYFMPKSTIYRGEMCFFDSED	PANSLRG	GEPNFLPV	TEEADI	REDDNIA	IIDV		
SEQ ID NO: 231	VGGACIYKYFMPKSTIYRGEMCFFDSED	PANSLRG	GEPNFLPV	TEEADI	REDDNIA	IIDV		
		40	50	60	70	80	90	
	130	140	150	160	170	180		
SEQ ID NO: 515	PVPSFSDSDPAAIIHDFEKGMTAYLD	LLLGNCY	LMPLNTS	IVMPPKN	LVELFG	KLASGRY		
SEQ ID NO: 231	PVPSFSDSDPAAIIHDFEKGMTAYLD	LLLGICY	LMPLNTS	IVMPPKN	LVELFG	KLASGRY		
		100	110	120	130	140	150	
	190	200	210	220	230	240		
SEQ ID NO: 515	LPQTYVVREDLVAVEEIRDVSNL	GIFIFYQ	LCNNRKS	FRLRRRD	LLLGFN	KRAIDKC	WKIR	
SEQ ID NO: 231	LPQTYVVREDLVAVEEIRDVSNL	GIFIFYQ	LCNNRKS	FRLRRRD	LLLGFN	KRAIDKC	WKIR	
		160	170	180	190	200	210	
	250	260						
SEQ ID NO: 515	HFPNEFIVETKICQE							
SEQ ID NO: 231	HFPNEFIVETKICQE							
		220						

**FIGURE 12**

[illegible]

```

          340          350
SEQ ID NO:196 AGTIYFLADLLVPTKARFP AFEL
              ::::::::::::::::::::::
SEQ ID NO:518 AGTIYFLADLLVPTKARFP AFEL
          380          390

```

**FIGURE 13**

98.5% identity in 194 aa overlap

```

          90      100      110      120      130      140
SEQ ID NO:519 ARNLPPPLTDAQKNKLRHLSVVTAAKVKCIPYAVLLEALALRNVRQLEDLVIEAVYADVL
               :
SEQ ID NO:158 ARNLPPPLTEAQKNKLRHLSVVTAAKVKCIPYAVLLEALALRNVRQLEDLVIEAVYADVL
          60      70      80      90      100      110

          150      160      170      180      190      200
SEQ ID NO:519 RGSLDQRNQRLEVVDYSIGRDIQRQDLSAIAQTLQEWCVGCEVVLSGIEEQVSRANQHKEQ
               :
SEQ ID NO:158 RGSLDQRNQRLEVVDYSIGRDIQRQDLSAIAQTLQEWCVGCEVVLSGIEEQVSRANQHKEQ
          120      130      140      150      160      170

          210      220      230      240      250      260
SEQ ID NO:519 QLGLKQQIESEVANLKKTIKVTAAAAAATSQDPEQHLTELREPASGTNQRQPSKKASKG
               :
SEQ ID NO:158 QLGLKQQIESEVANLKKTIKVTAAAAAATSQDPEQHLTELREPAPGTNQRQPSKKASKG
          180      190      200      210      220      230

          270
SEQ ID NO:519 KGLRGSAKIWSKSN
               :
SEQ ID NO:158 KGLRGSAKIWSKSN
          240      250

```

88.7% identity in 62 aa overlap

```

          10      20      30      40      50      60
SEQ ID NO:519 MSAEVKVTGQNQEQLLLAKSAKGAALATLIHQVLEAPGVYVFGELLDMPNVRELAESDF
               :
SEQ ID NO:158 MSAEVKVTGQNQEQLLLAKSAKGAALATLIHQVLEAPGVYVFGELLDMPNVRELXARNL
          10      20      30      40      50      60

SEQ ID NO:519 AS
               .X
SEQ ID NO:158 PP

```

**FIGURE 14**

68.9% identity in 74 aa overlap

```

              10      20      30      40      50
SEQ ID NO:226 MIARRNPVPLRFLPDEARSLPPPKLTDPRLLYIGFLGYCSGLIDNLIRRRPIATAGLHR
              .....
SEQ ID NO:514 MMTGRQGRATFQFLPDEARSLPPPKLTDPRLAFVGFGLGYCSGLIDNAIRRRPVLLAGLHR
              10      20      30      40      50      60

              60      70
SEQ ID NO:226 QLLYITAFFLLDIIL
              .....
SEQ ID NO:514 QLLYITSFVFVGYLLKRQDYMAYVRDHDMSYIKSHPEDFPEKDKKTYGEVFEEFHPVR
              70      80      90      100     110     120
```

FIGURE 15

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